

# robinson -FOOTPRINTING HYDROXYL RADICAL DNA

## => d his

```
(FILE 'HOME' ENTERED AT 11:55:51 ON 14 MAR 2002)
                SET COST OFF
     FILE 'REGISTRY' ENTERED AT 11:56:09 ON 14 MAR 2002
           3040 S .C.{2-4}C.{2-3}F....L..H...H/SQSP
L1
                SAV TEMP L1 HOPE424488/A
                E TGEK/SQEP
L2
              1 S E3
L3
              1 S E6
     FILE 'HCAPLUS' ENTERED AT 11:59:00 ON 14 MAR 2002
           1091 S L1
L4
              8 S L2 OR L3
L5
              6 S TGEK OR TGEKP
L6
              7 S L4 AND L5, L6
Ļ7
\Gamma8
            780 S L4 AND (ZN OR ZINC) (L) FINGER
             18 S L4 AND ALPHA(L) (HELICAL OR HELIX)
L9
L10
              3 S L4 AND QUAD?
            450 S L4 AND (DNA OR NUCLEIC ACID) (L) BINDING (L) PROTEIN
L11
             40 S L4 AND CYS2(L)HIS2
L12
              4 S L4 AND CYS()2(L)HIS()2
L13
             32 S L11 AND L12, L13
L14
            398 S L11 AND L8
L15
             11 S L15 AND L9, L10
L16
L17
             61 S L5-L7, L9, L10, L14, L16
                E WO98-53060/AP, PRN
                E W09853060/PN
              1 S E3
L18
                E WO98-GB1516/AP, RPN
                E W098-GB1516/AP, PRN
L19
              1 S E3, E4
L20
              1 S L18, L19
                E CHOO Y/AU
             71 S E3-E14
L21
                E KLUG A/AU
L22
            191 S E3, E4
                E ISALAN M/AU
             15 S E4
L23
             15 S L4 AND L18-L23
L24
             15 S L24 AND L8
L25
L26
             12 S L24 AND L11
L27
             6 S L24 AND L9, L10, L12, L13
              5 S L27 AND L26
L28
             6 S L27 AND L25
L29
            6 S L28, L29
L30
              5 S L30 NOT PROTON/TI
L31
             5 S L20,L31
L32
             10 S L24-L30 NOT L32
L33
             66 S L4 AND (DNA OR NUCLEIC ACID) (L) BINDING(L) (PEPTIDE OR POLYPEPT
L34
             45 S L34 AND L8
L35
             8 S L34 AND L12, L13
L36
         90 S L17, L32, L35, L36
L37
L38
             93 S L33, L37
            600 S L4 AND (PD<=19980526 OR PRD<=19980526 OR AD<=19980526)
L39
             59 S L39 AND L38
L40
                E DNA BINDING PROTEIN/CT
                E DNA-BINDING PROTEIN/CT
                E E4+ALL
           7172 S E1, E2, E3
L41
                E E2+ALL
```

52 S E3

L42



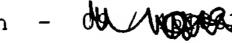
```
E DNA-BINDING PROTEIN/CT
                 E E4+ALL
                E E3+ALL
            108 S L41, L42 AND L39
L43
            266 S L39 AND L11, L34
L44
L45
            266 S L43, L44
            242 S L45 AND (ZN OR ZINC) (L) FINGER
L46
             23 S L46 AND L12, L13
L47
             33 S RECOGNI? AND L46
L48
                 E MOLECULAR RECOGNITION/CT
                 E E3+ALL
              7 S E2, E1+NT AND L46
L49
L50
               9 S E6+NT AND L46
              3 S RECOGNITION CODE AND L46
L51
L52
             35 S L47, L49-L51, L32
L53
             20 S L47, L48 NOT L52
L54
               6 S L53 AND (CONSEN? OR MOTIF)/TI
L55
               2 S L54 AND (BINDING PROTEIN)/TI
L56
             37 S L52, L55
             37 S L56 AND L4-L56
L57
     FILE 'HCAPLUS' ENTERED AT 12:25:10 ON 14 MAR 2002
     FILE 'BIOSIS' ENTERED AT 12:25:41 ON 14 MAR 2002
                 E CHOO Y/AU
             96 S E3-E12, E14
L58
                 E QUE L22
                 E KLUG A/AU
            155 S E3, E4
L59
                E ISALAN M/AU
               7 S E4
L60
               0 S L1
L61
L62
               0 S L2
L63
            237 S L58-L60
          49215 S (DNA OR NUCLEIC ACID) (L) BINDING(L) (PROTEIN OR PEPTIDE OR POLY
L64
             27 S L63 AND L64
L65
             16 S L65 AND PY<=1998
L66
               0 S L66 AND QUAD?
L67
               0 S L66 AND TETRA?
L68
L69
               1 S L66 AND TERT?
             13 S L66 AND (ZN OR ZINC)(L)FINGER
L70
L71
               3 S L66 NOT L70
               1 S L71 AND DESIGNING
L72
L73
             14 S L70, L72 AND L58-L72
     FILE 'HCAPLUS, BIOSIS' ENTERED AT 12:29:33 ON 14 MAR 2002
             49 DUP REM L57 L73 (2 DUPLICATES REMOVED)
L74
```

SET COST ON



#### => d his

(FILE 'HOME' ENTERED AT 11:55:51 ON 14 MAR 2002) Jan Delaval Reference Librarian SET COST OFF **Biotechnology & Chemical Library** FILE 'REGISTRY' ENTERED AT 11:56:09 ON 14 MAR 2002 CM1 1E07 - 703-308-4498  $3040 \text{ S} .C.\{2-4\}C.\{2-3\}F....L..H...H/SQSP$ jan.delaval@uspto.gov L1SAV TEMP L1 HOPE424488/A E TGEK/SQEP L21 S E3 L31 S E6 FILE 'HCAPLUS' ENTERED AT 11:59:00 ON 14 MAR 2002 1091 S L1 L4L58 S L2 OR L3 6 S TGEK OR TGEKP L6 7 S L4 AND L5, L6 L7 780 S L4 AND (ZN OR ZINC) (L) FINGER  $\Gamma8$ 18 S L4 AND ALPHA(L) (HELICAL OR HELIX) L93.S L4 AND QUAD? L10 450 S L4 AND (DNA OR NUCLEIC ACID) (L) BINDING (L) PROTEIN L1140 S L4 AND CYS2(L)HIS2 L124 S L4 AND CYS()2(L)HIS()2 L13 32 S L11 AND L12, L13 L14 L15 398 S L11 AND L8 11 S L15 AND L9, L10 L16 61 S L5-L7, L9, L10, L14, L16 L17 E W098-53060/AP, PRN E W09853060/PN L18 1 S E3 E WO98-GB1516/AP, RPN E W098-GB1516/AP, PRN L19 1 S E3, E4 L20 1 S L18, L19 E CHOO Y/AU L21 71 S E3-E14 E KLUG A/AU 191 S E3, E4 L22 E ISALAN M/AU L23 15 S E4 15 S L4 AND L18-L23 L24 15 S L24 AND L8 L25 12 S L24 AND L11 L26 L27 6 S L24 AND L9, L10, L12, L13 5 S L27 AND L26 L28 6 S L27 AND L25 L29 6 S L28, L29 L30 5 S L30 NOT PROTON/TI L31 L32 5 S L20, L31 10 S L24-L30 NOT L32 L33 66 S L4 AND (DNA OR NUCLEIC ACID) (L) BINDING(L) (PEPTIDE OR POLYPEPT L34 45 S L34 AND L8 L35 8 S L34 AND L12, L13 L36 90 S L17, L32, L35, L36 L37 93 S L33, L37 L38 600 S L4 AND (PD<=19980526 OR PRD<=19980526 OR AD<=19980526) L39 59 S L39 AND L38 L40 E DNA BINDING PROTEIN/CT E DNA-BINDING PROTEIN/CT E E4+ALL 7172 S E1, E2, E3 L41E E2+ALL L42 52 S E3 E DNA-BINDING PROTEIN/CT E E4+ALL E E3+ALL





```
108 S L41, L42 AND L39
L43
            266 S L39 AND L11, L34
L44
            266 S L43, L44
L45
            242 S L45 AND (ZN OR ZINC) (L) FINGER
L46
             23 S L46 AND L12, L13
L47
             33 S RECOGNI? AND L46
L48
                 E MOLECULAR RECOGNITION/CT
                 E E3+ALL
              7 S E2, E1+NT AND L46
L49
              9 S E6+NT AND L46
L50
L51
              3 S RECOGNITION CODE AND L46
             35 S L47, L49-L51, L32
L52
             20 S L47, L48 NOT L52
L53
              6 S L53 AND (CONSEN? OR MOTIF)/TI
L54
              2 S L54 AND (BINDING PROTEIN)/TI
L55
             37 S L52, L55
L56
             37 S L56 AND L4-L56
L57
```

=> fil hcaplus FILE 'HCAPLUS' ENTERED AT 12:25:10 ON 14 MAR 2002 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 14 Mar 2002 VOL 136 ISS 11 FILE LAST UPDATED: 12 Mar 2002 (20020312/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

The P indicator for Preparations was not generated for all of the CAS Registry Numbers that were added to the CAS files between 12/27/01 and 1/23/02. As of 1/23/02, the situation has been resolved. Searches and/or SDIs in the H/Z/CA/CAplus files incorporating CAS Registry Numbers with the P indicator executed between 12/27/01 and 1/23/02 may be incomplete. See the NEWS message on this topic for more information.

=> d 157 bib abs hitrn retable tot

- ANSWER 1 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57
- 2002:51500 HCAPLUS ΑN
- 136:114558 DN
- Zinc finger synthetic polypeptides TIbinding to telomeric G quadruplex DNA
- Choo, Yen; Isalan, Mark; Liu, Xiaohai; Patel, Sachin; INBalasubramanian, Shankar
- Sangamo Biosciences, Inc., USA; Cambridge University Technical Services PΑ Ltd.
- PCT Int. Appl., 147 pp. SO CODEN: PIXXD2





```
Patent
DT
     English
LA
FAN.CNT 1
                                           APPLICATION NO.
     PATENT NO.
                      KIND DATE
                                                            DATE
PΙ
                      Α2
                            20020117
                                           WO 2001-GB3130
                                                            20010712
     WO 2002004488
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
             RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
             UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                            20000712
PRAI US 2000-614679
                      Α
     Nucleic acid-binding polypeptides
AB
     are designed capable of binding to one or more of telomeric, G-
     quadruplex, or G-quartet nucleic acid as an
     inhibitor of enzymic activity, including a telomerase activity, a
     polymerase activity, an integrase activity, and a gp120 activity. Gq1 is
     an artificial protein that has been engineered from zinc
     finger motifs to bind human telomeric G-quadruplex
     DNA. Primer extension studies using both telomerase and Klenow
     fragment of Escherichia coli DNA polymerase I suggest that Gq1
     can inhibit both the synthesis and copying of telomeric DNA
     sequences. Since this zinc finger protein
     has no detectable affinity for telomeric duplex DNA, Gql may
     prove an attractive probe for carrying out cell-based studies. Telomerase
     assays as well as methods of identifying mols. capable of interacting with
     telomeric, G-quadruplex, or G-quartet nucleic
     acid are described. A stable integrated Gql zinc
     finger repressor is shown to inhibit HIV-1 replication in human
     T-cells.
     390883-06-4P 390883-09-7P 390883-11-1P
IT
     390883-13-3P 390883-15-5P
     RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
     PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (amino acid sequence; encoding zinc finger
        synthetic polypeptides binding to telomeric G
       quadruplex DNA)
IT
     390883-30-4
    RL: PRP (Properties)
        (unclaimed sequence; zinc finger synthetic
       polypeptides binding to telomeric G
       quadruplex DNA)
    ANSWER 2 OF 37 HCAPLUS COPYRIGHT 2002 ACS
L57
AN
     2001:408061 HCAPLUS
     135:30537
DN
     Design, construction and of zinc finger protein
TI
     derivatives and their use in the modulation of gene expression
     Barbas, Carlos F., III; Gottesfeld, Joel M.; Wright, Peter E.
IN
     The Scripps Research Institute, USA
PA
    U.S., 67 pp., Cont.-in-part of U.S. Ser. No. 312,604, abandoned.
SO
     CODEN: USXXAM
DT
     Patent
    English
LA
FAN.CNT 2
     PATENT NO.
                                           APPLICATION NO.
                     KIND DATE
                                                            DATE
                    B1 20010605
    US 6242568
                                           US 1996-676318 19961230 <--
PI
                                           WO 1995-US829 19950118 <--
    WO 9519431 A1 19950720
```

W: AU, CA, FI, JP, NO, US





RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE B2 19940118 <--PRAI US 1994-183119 US 1994-312604 B2 19940928 <--19950118 <--WO 1995-US829 The present invention provides zinc finger nucleotide AB binding protein variants that have at least two zinc finger modules that bind to a target cellular nucleotide sequence and modulate the transcriptional function of the cellular nucleotide sequence. Also provided are methods of use of such zinc finger nucleotide binding protein variants and methods for isolating the same using expression libraries encoding the protein variants contg. randomized substitutions of amino acids. Exemplary zinc finger nucleotide binding protein variants of the invention include two cysteines and two histidines whereby both cysteines are amino proximal to both histidines. Design and construction of variants of the zinc finger protein Zif/268 are disclosed. Construction of multifinger proteins utilizing repeats of the first finger of Zif/268 is described. 343429-13-0P IT RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BPR (Biological process); PRP (Properties); BIOL (Biological study); PREP (Preparation); PROC (Process) (amino acid sequence; design, construction and of zinc finger protein derivs. and their use in modulation of gene expression) 169108-70-7P 169108-73-0P 169108-74-1P IT 169108-76-3P RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BPR (Biological process); PRP (Properties); BIOL (Biological study); PREP (Preparation); PROC (Process) (design, construction and of zinc finger protein derivs. and their use in modulation of gene expression) 343581-44-2 IT RL: PRP (Properties) (unclaimed protein sequence; design, construction and of zinc finger protein derivs. and their use in the modulation of gene expression) 343321-37-9 ITRL: PRP (Properties) (unclaimed sequence; design, construction and of zinc finger protein derivs. and their use in the modulation of gene expression) 168971-84-4 IT RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological study); USES (Uses) (zinc finger linker peptide; design, construction and of zinc finger protein derivs. and their use in modulation of gene expression) RETABLE | Referenced Work | Referenced Referenced Author |Year | VOL | PG File | (RPY) | (RVL) | (RPG) (RWK) (RAU) |Biochemistry | HCAPLUS |1991 |30 |7842 Agarwal, K |Proceedings of the N|HCAPLUS |1992 |89 | 4457 Barbas, C | Nucleic Acids Resear | HCAPLUS 12715 |1990 |18 Bergqvist, A IUS 5350840 | HCAPLUS 11994 | Call |1986 |233 |Science 11175 | HCAPLUS Celenza, J IUS 5376530 | HCAPLUS |1994 | de The |10189 | The Journal of Biolo | HCAPLUS |1990 |265 Debs, R IUS 5597693 HCAPLUS |1997 | Evans US 5243041 | HCAPLUS 11993 Fernandez-Pol |The EMBO Journal | HCAPLUS |1992 |11 14507 Jacobs, G

|5689

143

|1994 |33

|1991 |

|1992 |

|1993 |331

Jamieson, A

Julian, N

Katagiri

Ladner

| Biochemistry

|FEBS Letters

IUS 4990607

|US 5096815

HCAPLUS

| HCAPLUS

| HCAPLUS

| HCAPLUS





Ladner Nabel Pabo, C Quigley, C Rauscher, F Ray, A Rollins, M Singh, H South, T Stevens Tao	1988  52  1990  29  1994    1994	1103  1259  7086  4776  415  7786	US 5403484
		  2784  588  6340	US 5324638   HCAPLUS   Molecular and Cellul   MEDLINE

- L57 ANSWER 3 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1999:686615 HCAPLUS
- DN 131:308157
- TI cDNA molecules encoding human and chicken CTCF (CCCTC-binding factor), sequences and uses thereof
- IN Lobanenkov, Victor L.; Neiman, Paul E.; Klenova, Elena M.; Goodwin, Graham H.; Filippova, Galina N.; Collins, Steven J.; Nicolas, Robert H.
- PA Fred Hutchinson Cancer Research Center, USA; Cancer Research Campaign Technology, Ltd.
- SO U.S., 37 pp., Cont.-in-part of U.S. Ser. No. 261,680, abandoned. CODEN: USXXAM
- DT Patent
- LA English
- FAN.CNT 2

	PATENT NO.	KIND	DATE		APPLICATION NO.	DATE
ΡI	US 5972643	Α	19991026		US 1995-475844	19950607 <
	CA 2205203	AA	19951228		CA 1995-2205203	19950615 <
PRAI	US 1994-261680		19940617	<		

AB This invention provides the isolation and purifn. of polynucleotides (genomic DNA, cDNA, antisense RNA) encoding human and chicken CCCTC-binding factor (CTCF). CTCF is a sequence-specific DNA binding protein that contains eleven zinc finger binding domains and is capable of binding to the 5'-flanking region of the c-myc gene. The invention also provides methods for producing recombinant CTCF by inserting nucleic acid mols. encoding CTCF into a suitable expression vector and use of said expression vector to transform prokaryotic or enkaryotic cells. The cDNA and amino acid sequences of

prokaryotic or eukaryotic cells. The cDNA and amino acid sequences of chicken and human CTCF are provided. Using Northern blot anal. the invention revealed four major chicken CTCF gene mRNA species, indicating that the CTCF gene may encode multiple proteins by generating a variety of mRNA isoforms. The invention further showed that CTCF acts as a repressor of the human c-myc gene P2 promoter. The invention also discussed the potential use of CTCF polypeptides and antibodies to identify mutant CTCFs in methods of diagnosis.

IT 152890-29-4P 177404-57-8P

RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BPR (Biological process); PRP (Properties); BIOL (Biological study); PREP (Preparation); PROC (Process) (amino acid sequence; human and chicken CTCF (CCCTC-binding factor), cDNA and amino acid sequences, recombinant prodn. and binding to gene c-myc 5'-flanking region)

### RETABLE

Referenced Author (RAU)	Year   VOL   PG  (RPY) (RVL) (RPG)	Referenced Work   Referenced   (RWK)   File
=======================================	=+====+======	=+=====================================
Arnold	1996  24  2640	Nucleic Acids Resear HCAPLUS
Askew	1991  6  1915	Oncogene   HCAPLUS
Basu	1993  268  4188	J Biol Chem   HCAPLUS
Ben-David	1991  66  831	Cell  MEDLINE





					UCA DI IIC
Ben-David	•		1332	Proc Natl Acad Sci U	
Berg	1990	265	6513	[ • — — — — ·	HCAPLUS
Beug	•	1	195	J Cell Physiol Suppl	WEDLINE
Bickmore	11992	257 <b> </b>	235		HCAPLUS
Bird	1986	321	209	Nature	HCAPLUS
Blackwood	1991	251	1211	,002000	HCAPLUS
Bossone	1992	89	7452	,	HCAPLUS
Burcin	1994	5 1	337	100::002 =====51	HCAPLUS
Carter	•	87 I	8751	Proc Natl Acad Sci U	MEDLINE
Cole	•	20	361	Annu Rev Genet	HCAPLUS
El-Baradi	•	35	155	Mech Devel	HCAPLUS
Flanagan	1992	12	38	Mol Cell Biol	MEDLINE
Franklin	1994	114	6773	Mol Cell Biol	HCAPLUS
	1981	19	6505		<b>HCAPLUS</b>
Fried	1989	186	1934		<b>HCAPLUS</b>
Gould	1992	170	337	•	<b>HCAPLUS</b>
Helin	11992	257	1946	Science	HCAPLUS
Hsu	11985	1313	1806	Nature	HCAPLUS
Jacobs	•	•	110	Methods Enzymol	HCAPLUS
Kadonaga	11991	•	1566	Science	HCAPLUS
Kadonaga	11988	•	•	Mol Cell Biol	HCAPLUS
Kim	1990	110	3224	Mol Cell Biol	HCAPLUS
Kingsley	11992	12	4251	•	HCAPLUS
Klenova	11993	•	7612	Mol Cell Biol	HCAPLUS
Kohlhuber	11993	18	11099	Oncogene	MEDLINE
Kohne	11993	232	1747	J Mol Biol	I IMEDTINE
Kolluri	1991	117	14771	Nucl Acids Res	 
Kretzner	1992	359	1426	Nature	HCAPLUS
Krumm	1992	16	2201	Genes 7 Devel	HCAPLUS
Lathe	1985	1183	1	J Mol Biol	HCAPLUS
Lobanenkov	1986	159	181	Eur J Biochem	HCAPLUS
Lobanenkov	1989		45	Gene Reg and AIDS	
Lobanenkov	1990	15	1743	Oncogene	HCAPLUS
Marcu	1992	61	1809		HCAPLUS
Matsudaira	11987	1262	10035	J Biol Chem	HCAPLUS
Mattes	11992	1		US 5143843	HCAPLUS
Morishita	11992	189	3937	1	HCAPLUS
Neiman	11991	188	5857	Proc Nal Acad Sci US	
Ngo	1994		1433	The Protein Folding	HCAPLUS
Nicolas	11993	1	81	Transcription Factor	HCAPLUS
Pyrc	11992	31	4102	Biochem	HCAPLUS
Ray	11991	11	2154	Mol Cell Biol	HCAPLUS
Riggs	1993	113	17487	Mol Cell Biol	HCAPLUS
Roy	11991	1354	245	Nature	HCAPLUS
Sambrook	1989	İ	8.2	Molecular Cloning: A	•
Sen	11986	146	1705	Cell	HCAPLUS
Shih	1984	181	14697	Proc Natl Acad Sci U	HCAPLUS
Smulson	1993	İ	i	US 5272057	HCAPLUS
Spencer	1991	156	11	Adv Cancer REs	HCAPLUS
St-Arnaud	11993	113	1590	Mol Cell Biol	HCAPLUS
	11992	120	624	Nucl Acids Res	HCAPLUS
Stappert	11992	111	3307	EMBO J	1
Stobl	11991	125	11013	Molecul Biol (Moscow	MEDLINE
Tevosian	11990	187	6791	Proc Natl Acad Sci U	HCAPLUS
Tsuda van Lohuizen	11990	1032	1213	Biochimica et Biophy	
vaii monutaen	1200	,	,	· •	

- · L57 ANSWER 4 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1999:614102 HCAPLUS
- DN 131:238849
- TI A DNA-binding zinc finger
  protein specific for a modified base-containing sequence
- IN Choo, Yen; Isalan, Mark
- PA Medical Research Council, UK
- SO PCT Int. Appl., 56 pp. CODEN: PIXXD2
- DT Patent
- LA English





```
FAN.CNT 1
                                           APPLICATION NO.
                                                            DATE
                      KIND
                            DATE
     PATENT NO.
                                           WO 1999-GB816
                                                            19990317 <--
                            19990923
                       A2
    WO 9947656
ΡI
                            19991125
    WO 9947656
                       А3
            AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
             DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
             JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
             MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
             TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
             MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                      AU 1999-29449
                                                            19990317 <--
                           19991011
    AU 9929449
                       A1
                                           EP 1999-910512
                                                            19990317 <--
                            20010103
                       A2
     EP 1064369
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
                                                            19990317 <--
                                           JP 2000-536839
                            20020305
                       Т2
     JP 2002506640
                            19980317
                                      <--
PRAI GB 1998-5576
                       Α
                            19980331
                                      <--
     GB 1998-6895
                       A
                                     <--
                            19980403
                       Α
     GB 1998-7246
     WO 1999-GB816
                       W
                            19990317
     A modified Cys2-His2 zinc finger
AB
     that binds to a target nucleic acid sequence contg. a modified base but
     not to an identical sequence contg. an equiv. unmodified base is
     described. The zinc finger can be used to create
     sequence-specific reagents, such as restriction enzymes with novel
     sequence requirements or for the assay of DNA methylation. Zinc
     fingers capable of binding 5-Me cytosine-contg. DNA were derived
     from one of the fingers of Zif268 (Egr-1) by several rounds of
     screening of a phage display library using increasing stringency of
     selectivity of binding to screen for sequence-specific, selective binding.
     Rules relating the amino acid sequence of a zinc finger
     to its DNA binding specificity are also outlined. Zinc
     fingers showing up to a 31-fold difference in dissocn. consts. of
     <100 nM for binding with 5-Me cytosine-contg. DNA and its cytosine-contg.
     analog were obtained.
     133551-05-0 216434-95-6
IT
     RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
        (methylated DNA binding zinc
        finger; DNA-binding zinc
        finger protein specific for modified base-contg.
        sequence)
     ANSWER 5 OF 37 HCAPLUS COPYRIGHT 2002 ACS
L57
     1998:790665 HCAPLUS
AN
     130:35366
DN
     Recognition code for the design of synthetic
TI
     nucleic acid-binding proteins
     Choo, Yen; Klug, Aaron; Isalan, Mark
IN
     Medical Research Council, UK
PA
     PCT Int. Appl., 57 pp.
SO
     CODEN: PIXXD2
     Patent
\mathsf{DT}
     English
LA
FAN.CNT 3
                                           APPLICATION NO.
                                                           DATE
                      KIND DATE
     PATENT NO.
                                                             19980526 <--
                                            WO 1998-GB1516
                             19981126
                      A1
     WO 9853060
ΡI
         W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
             UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
```



```
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
            FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
            CM, GA, GN, ML, MR, NE, SN, TD, TG
                                         AU 1998-75426
                                                          19980526 <--
                          19981211
                      A1
    AU 9875426
                           20010412
                      B2
    AU 732017
                                          EP 1998-922967
                                                          19980526 <--
                           20000308
                      A1
    EP 983351
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, FI
                                                          19980526 <--
                                          JP 1998-550158
                      Т2
                           20011225
    JP 2001527417
                           19970523
PRAI GB 1997-10809
                      Α
                           19980526 <--
    WO 1998-GB1516
    MARPAT 130:35366
OS
    The invention provides a method for prepg. a nucleic
AB
    acid binding protein of the Cys2-
    His2 zinc finger class capable of
    binding to a target quadruplet nucleic
    acid sequence. A more complete code is provided which permits the
     selection of any nucleic acid sequence as the target
     sequence, and the design of a specific nucleic acid-
    binding protein which will bind thereto. Moreover, the
     invention provides a method by which a zinc finger
    protein specific for any given nucleic acid
     sequence may be designed and optimized. If bas 4 in the
     quadruplet is A, then position +6 in the .alpha.-
    helix is Gln and position ++2 is not Asp; and if base 4 in the
     quadruplet is C, then position +6 in the .alpha.-
     helix may be any residue, as long as position ++2 in the .
     alpha.-helix is not Asp. The recognition
     code is used to design (1) a protein whereby the target
     is the activating point mutation in codon 12 of the human EJ bladder
     carcinoma Ha-ras oncogene (GGC.fwdarw.GTC), and (2) an anti-HIV
     zinc finger binding to the tat-specific
     sequence 5'-agagagctc-3'.
     216493-25-3P
IT
     RL: BPN (Biosynthetic preparation); BPR (Biological process); PRP
     (Properties); BIOL (Biological study); PREP (Preparation); PROC (Process)
        (anti-HIV zinc finger designed for specific
        nucleic acid binding; recognition
        code for the design of synthetic nucleic acid
        -binding proteins)
     216434-95-6P
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); BIOL (Biological
     study); PREP (Preparation)
        (model consensus zinc finger; recognition
        code for the design of synthetic nucleic acid
        -binding proteins)
     216434-99-0P 216435-00-6P 216435-01-7P
IT
     216437-56-8P 216437-57-9P 216437-58-0P
     216437-59-1P 216437-60-4P 216437-61-5P
     216582-15-9P 216583-28-7P
     RL: BPN (Biosynthetic preparation); PRP (Properties); BIOL (Biological
     study); PREP (Preparation)
        (zinc finger designed for binding to
        Ha-ras oncogene; recognition code for the design of
        synthetic nucleic acid-binding
        proteins)
RETABLE
                                                              | Referenced
                                          | Referenced Work
                       |Year | VOL | PG
   Referenced Author
                                                               | File
                       |(RPY)|(RVL)|(RPG) |
                                                 (RWK)
         (RAU)
|WO 9606166 A
                                                               | HCAPLUS
                       |1996 |
Choo, Y
                                          [Curr Op Struct Biol | HCAPLUS
                       |1997 |7
                                   |117
Choo, Y
                                   |11163 | Proceedings of the N| HCAPLUS
                       |1994 |91
Choo, Y
                                   |11168 | Proceedings of the N| HCAPLUS
                       |1994 |91
Choo, Y
                                   |1171 |Structure
                                                              HCAPLUS
                       |1996 |4
Elrod-Erickson, M
                                          |Proceedings of the N|HCAPLUS
                                   |5617
                       11997 | 94
```

Isalan, M

```
robinson - 09 / L57 ANSWER 6 OF 37 HCAPLUS COPYRIGHT 2002 ACS
AN 1998:790664 HCAPLUS
DN 130:35365
TI Recognition code for the design of synthetic nucleic acid-binding proteins
```

```
ΑN
     Choo, Yen; Klug, Aaron; Isalan, Mark
IN
PA
     Medical Research Council, UK
     PCT Int. Appl., 63 pp.
SO
     CODEN: PIXXD2
     Patent
DT
     English
LA
FAN.CNT 1
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                            DATE
                            19981126
                                                            19980526 <--
                                           WO 1998-GB1514
PI
    WO 9853059
                       A1
            AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
             UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
             CM, GA, GN, ML, MR, NE, SN, TD, TG
                                                            19980526 <--
                                           AU 1998-75424
                       A1
    AU 9875424
                            19981211
PRAI GB 1997-10807
                            19970523 <--
                            19980526 <--
     WO 1998-GB1514
OS
    MARPAT 130:35365
    The invention provides a method for prepg. a nucleic
AB
    acid binding protein of the Cys2-
    His2 zinc finger class capable of
    binding to a target triplet nucleic acid
    sequence. A more complete code is provided which permits the selection of
    any nucleic acid sequence as the target sequence, and
    the design of a specific nucleic acid-binding
    protein which will bind thereto. Moreover, the invention provides
    a method by which a zinc finger protein
    specific for any given nucleic acid sequence may be
    designed and optimized. Binding to the 5' base of the triplet
    by an .alpha.-helical zinc finger
    nucleic acid binding motif in the
    protein is detd. as follows: if the 5' base in the triplet is A,
    then position +6 in the .alpha.-helix is Glu, Asn or
    Val; if the 5' base in the triplet is C, then position +6 in the .
    alpha.-helix is Ser, Thr, Val, Ala, Glu or Asn. The
    recognition code is used to design (1) a protein
    whereby the target is the activating point mutation in codon 12 of the
    human EJ bladder carcinoma Ha-ras oncogene (GGC.fwdarw.GTC), (2) an
    anti-HIV zinc finger binding to the
    tat-specific sequence 5'-agagagete-3', and (3) design of a zinc
    finger specific for an 8-bp palindrome (gcggccgc).
    216493-25-3P
IT
    RL: BPN (Biosynthetic preparation); BPR (Biological process); PRP
     (Properties); BIOL (Biological study); PREP (Preparation); PROC (Process)
        (anti-HIV zinc finger designed for specific
       nucleic acid binding; recognition
        code for the design of synthetic nucleic acid
        -binding proteins)
IT
    216434-95-6P
    RL: BPN (Biosynthetic preparation); PRP (Properties); BIOL (Biological
    study); PREP (Preparation)
        (model consensus zinc finger; recognition
        code for the design of synthetic nucleic acid
        -binding proteins)
    216434-99-0P 216435-00-6P 216435-01-7P
IT
```

216582-68-2P



L57

AN

DN

TI

IN

PA

SO

DT

LA

PI

OS

AΒ



RL: BPN (Biosynthetic preparation); PRP (Properties); BIOL (Biological study); PREP (Preparation) (zinc finger designed for binding to Ha-ras oncogene; recognition code for the design of synthetic nucleic acid-binding proteins) RETABLE |Year | VOL | PG | Referenced Work | Referenced Referenced Author I File |(RPY)|(RVL)|(RPG) | (RWK) (RAU) IWO 9606166 A HCAPLUS |1996 | Choo, Y |Curr Op Struct Biol | HCAPLUS |1997 |7 |117 Choo, Y |11163 | Proceedings of the N| HCAPLUS |1994 |91 Choo, Y |11168 | Proceedings of the N| HCAPLUS |1994 |91 Choo, Y |1171 |Structure | HCAPLUS |1996 |4 Elrod-Erickson, M HCAPLUS 1483 | Nature |1993 |366 Fairall, L |13577 | Proceedings of the N| HCAPLUS |1996 |93 Houbaviy, H |MEDLINE |1996 |181 |167 | |Gene Ikeda, M |5617 | Proceedings of the N| HCAPLUS |1997 |94 Isalan, M |5689 |Biochemistry | HCAPLUS 11994 | 33 Jamieson, A |1701 |Science HCAPLUS |1993 |261 Pavletich, N |12357 | Proceedings of the N| HCAPLUS |1994 |91 Suzuki, M | Proc Natl Acad Sci | HCAPLUS 344 11995 192 Wu, H ANSWER 7 OF 37 HCAPLUS COPYRIGHT 2002 ACS 1998:790663 HCAPLUS 130:35364 Recognition code for the design of synthetic nucleic acid-binding proteins Choo, Yen; Klug, Aaron; Isalan, Mark Medical Research Council, UK PCT Int. Appl., 63 pp. CODEN: PIXXD2 Patent English FAN.CNT 3 APPLICATION NO. DATE KIND PATENT NO. WO 1998-GB1512 19980526 <--19981126 A1 WO 9853058 W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG AU 1998-75423 19980526 <--19981211 A1AU 9875423 EP 1998-922964 19980526 <--20000308 **A**1 EP 983350 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI 19970523 <---PRAI GB 1997-10809 Α W · 19980526 <--WO 1998-GB1512 MARPAT 130:35364 The invention provides a method for prepg. a nucleic acid binding protein of the Cys2-His2 zinc finger class capable of binding to a target quadruplet nucleic acid sequence. Zinc finger binding sites are detd. by overlapping 4-bp subsites, and sequence specificity at the boundary between subsites arises from synergy between adjacent finger.s. A more complete code is provided which permits the selection of any nucleic acid sequence as the target sequence, and the design of a specific nucleic acid-

binding protein which will bind thereto. Moreover, the

invention provides a method by which a zinc finger





protein specific for any given nucleic acid sequence may be designed and optimized. Binding to base 4 of the quadruplet by an .alpha.-helical zinc finger nucleic acidbinding motif in the protein is detd. as follows: if base 4 in the quadruplet is A, then position +6 in the . alpha.-helix is Glu, Asn, or Val; if base 4 in the quadruplet is C, then position +6 in the .alpha.helix is Ser, Thr, Val, Ala, Glu, or Asn. The recognition code is used to design (1) a protein whereby the target is the activating point mutation in codon 12 of the human EJ bladder carcinoma Ha-ras oncogene (GGC.fwdarw.GTC), (2) an anti-HIV zinc finger binding to the tat-specific sequence 5'-agagagete-3', and (3) design of a zinc finger specific for an 8-bp palindrome (gcggccgc). 216493-25-3P IT RL: BPN (Biosynthetic preparation); BPR (Biological process); PRP (Properties); BIOL (Biological study); PREP (Preparation); PROC (Process) (anti-HIV zinc finger designed for specific nucleic acid binding; recognition code for the design of synthetic nucleic acid -binding proteins) 216434-95-6P IT RL: BPN (Biosynthetic preparation); PRP (Properties); BIOL (Biological study); PREP (Preparation) (model consensus zinc finger; recognition code for the design of synthetic nucleic acid -binding proteins) 216434-99-0P 216435-00-6P 216435-01-7P IT 216582-68-2P RL: BPN (Biosynthetic preparation); PRP (Properties); BIOL (Biological study); PREP (Preparation) (zinc finger designed for binding to Ha-ras oncogene; recognition code for the design of synthetic nucleic acid-binding proteins) RETABLE | Referenced Work | Referenced |Year | VOL | PG Referenced Author |(RPY)|(RVL)|(RPG)| | File (RWK) HCAPLUS IWO 9606166 A 11996 I |5617 | Proceedings of the N| HCAPLUS |1997 |94 Isalan, M ANSWER 8 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57 1998:786691 HCAPLUS AN DN130:149362 Isolation and functional characterization of cDNA of serum amyloid TIA-activating factor that binds to the serum amyloid A promoter Ray, Alpana; Ray, Bimal K. ΑU Department of Veterinary Pathobiology, University of Missouri, Columbia, CS MO, 65211, USA Mol. Cell. Biol. (1998), 18(12), 7327-7335 SO CODEN: MCEBD4; ISSN: 0270-7306 American Society for Microbiology PBJournal  $\mathsf{DT}$ English LASerum amyloid A (SAA), a plasma protein inducible in response to AΒ many inflammatory conditions, is assocd. with the pathogenesis of several diseases including reactive amyloidosis, rheumatoid arthritis, and atherosclerosis. The authors have previously reported an element of the SAA promoter, designated SAA-activating sequence (SAS), that is involved in the inflammation-induced SAA expression, and a nuclear factor, SASbinding factor (SAF), that interacts with the SAS element has been identified previously (A. Ray and B. K. Ray, Mol. Cell. Biol. 16:1584-1594, 1996). To evaluate how SAF is involved in SAA promoter

activation, the authors have investigated structural features and

robinson - 09 / 000

functional characteristics of this transcription factor. These studies indicate that SAF belongs to a family of transcription factors characterized by the presence of multiple zinc finger motifs of the Cys2-His2 type at the carboxyl end. Of the three cloned SAF cDNAs (SAF-1, SAF-5, and SAF-8), SAF-1 isoform showed a high degree of homol. to MAZ/ZF87/Pur-1 protein while SAF-5 and SAF-8 isoforms are unique and are related to SAF-1/MAZ/ZF87/Pur-1 at the zinc finger domains but different elsewhere. Although structurally distinct, all members are capable of activating SAS element-mediated expression and display virtually identical sequence specificities. However, varying levels of expression of members of this gene family were obsd. in different tissues. Functional activity of SAF is regulated by a posttranslational event as SAF DNAbinding and transactivation abilities are increased by a protein phosphatase inhibitor, okadaic acid, and inhibited by a protein kinase inhibitor, H7. Consistent with this observation, increased DNA binding of the cloned SAF and its hyper-phosphorylation, in response to okadaic acid treatment of the transfected cells, were obsd. Taken together, our results suggest that, in addn. to tissue-specific expression, SAFs, a family of zinc finger transcription factors, undergo a modification by a posttranslational event that confers their SAA promoter-binding activity and transactivation potential.

220202-75-5 220202-76-6 220202-78-8 IT

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL

(Biological study)

(amino acid sequence; isolation and functional characterization of cDNA of serum amyloid A-activating factor that binds to serum amyloid promoter)

(RAU)	Year  (RPY)	(RVL)	(RPG)	Referenced Work   (RWK)	Referenced   File
(RAU)  ===================================	+====  1995  1991  1998  1997  1998  1998  1998  1998  1998  1998  1998  1986  1986	====================================	======   3110   15277   211   36   25624   7452   248   509   273   156   9644   3179   1908   456   4475   1701	Proc Natl Acad Sci U   J Biol Chem   Cell   Arthritis Rheum   J Biol Chem   Proc Natl Acad Sci U   Anal Biochem   Cell   Structure and functi   Anal Biochem   Cell Regul   J Biol Chem   Mol Cell Biol   Mol Cell Biol   Virology   Mol Cell Biol   Froc Natl Acad Sci U   Cell   Nucleic Acids Res   Ann N Y Acad Sci	HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS HCAPLUS
Poole, S Pyrc, J Ray, A	•	31  3	4102  151	Biochemistry  Gene Expr	HCAPLUS   HCAPLUS

	11005	1270	17365	J Biol Chem	HCAPLUS
Ray, A	11995		•	•	•
Ray, A	1994	14	4324	Mol Cell Biol	HCAPLUS
Ray, A	11996	116	1584	Mol Cell Biol	HCAPLUS
Ray, B	11992	1185	169	Biochem Biophys Res	HCAPLUS
Ray, B	11993	1193	11159	Biochem Biophys Res	HCAPLUS
	11997	136	14662	Biochemistry	HCAPLUS
<b>4</b> •	11997	1272	•	J Biol Chem	HCAPLUS
Ray, B	11991	1266	120270	J Biol Chem	HCAPLUS
Rossomando, A	•	1 2 0 0	1	Molecular cloning:a	İ
Sambrook, J	11989	!	1	•	HCAPLUS
Singh, H	1988	52	415	[Cell	,
Sipe, J	11992	161	1947	Annu Rev Biochem	HCAPLUS
Steinmetz, A	11989	1006	173	Biochim Biophys Acta	HCAPLUS
Sturgill, T	11991	11092	1350	Biochim Biophys Acta	
Wadzinski, B	1993	i13	12822	Mol Cell Biol	HCAPLUS
Wegenka, U	1993	113	1276	Mol Cell Biol	HCAPLUS
Williamson, M	11994	1297	1249	Biochem J	HCAPLUS
•	•	110	6181	Mol Cell Biol	HCAPLUS
Wilson, D	11990	• —	•	•	HCAPLUS
Zhang, D	1996	1271	9503	J Biol Chem	LUCATION
			_		

L57 ANSWER 9 OF 37 HCAPLUS COPYRIGHT 2002 ACS

AN 1998:704604 HCAPLUS

DN 130:61946

TI Cloning the cDNA for a new human zinc finger protein defines a group of closely related Kruppel-like transcription factors

AU Matsumoto, Nobukyuki; Laub, Friedrich; Aldabe, Rafael; Zhang, Wen; Ramirez, Francesco; Yoshida, Teruhiko; Terada, Masaaki

CS Genetics Division, National Cancer Center Research Institute, Tokyo, 104-0045, Japan

SO J. Biol. Chem. (1998), 273(43), 28229-28237 CODEN: JBCHA3; ISSN: 0021-9258

PB American Society for Biochemistry and Molecular Biology

DT Journal

LA English

We have identified a novel zinc finger protein
that has been named ubiquitous Kruppel-like factor (UKLF) based on
structural considerations and the pattern of gene expression. UKLF was
isolated by the polymerase chain reaction approach using degenerate
oligonucleotides corresponding to the DNA-binding
domain of erythroid Kruppel-like factor (EKLF) and cDNA prepd. from human
vascular endothelial cells. The carboxyl-terminal portion of UKLF
contains three zinc fingers of the Cys2His2 type and binds in vitro to the CACCC motif of the
beta,-globin promoter and to the Sp1 recognition sequence. The

beta.-globin promoter and to the Sp1 recognition sequence. The amino-terminal portion of UKLF consists of a hydrophobic region rich in serines and a neg. charged segment with several glutamic acid residues. The first 47 amino acids of the acidic region are nearly identical to the amino-terminal portion of another Kruppel-like factor, the so-called core promoter-binding protein (CPBP) or Zf9. Like

promoter-binding protein (CPBP) or ZI9. Like CPBP/Zf9, UKLF can function as a transcription activator in co-transfection assays. However, this activity is lost when the highly conserved amino-terminal segment is deleted. These findings indicate that UKLF and CPBP/Zf9 represent a distinct subgroup of closely related Kruppel-like activators of transcription. Mapping of the UKLF gene to chromosome 2 suggested that UKLF and CPBP/Zf9 translocated to different chromosomes following duplication from an ancestral gene.

IT **217798-42-0** 

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); PRP (Properties); BIOL (Biological study); PROC (Process) (amino acid sequence; cloning the cDNA for a new human zinc finger protein defines a group of closely related Kruppel-like transcription factors)

RETABLE Referenced Author (RAU)	Year   VOL   P  (RPY) (RVL) (R	RPG)   (RWK)	Referenced   File
Anderson, K	·	====+=================================	HCAPLUS



	robinson		,
1990  265	16513	J Biol Chem	HCAPLUS
11996   15	5888	EMBO J	HCAPLUS
11987   162	•	Anal Biochem	HCAPLUS
11996   16	1695	Mol Cell Biol	HCAPLUS
11996   13	2623	Oncogene	HCAPLUS
1996   14	1101	Prog Liver Dis	HCAPLUS
11996   271	31384	J Biol Chem	1
1987  51	1121	Cell	HCAPLUS
11993  72	1481	Cell	HCAPLUS
14000 100	10500	IDaga Nati Agad Cai	THECAPTIIS

Hahn, S |Proc Natl Acad Sci U|HCAPLUS |1992 |89 19509 Heng, H |HCAPLUS |Biochem J |1995 |311 1675 Ide, H | HCAPLUS IJ Biol Chem |14828 1269 11994 Inagaki, Y HCAPLUS

|J Clin Invest 187 1847 |1991 Johnson, R 11367 |Science |1991 |253 Klevitt, R | HCAPLUS |Eur J Biochem |1996 |236 1365 Koritschoner, N

HCAPLUS 11997 |272 19573 | J Biol Chem Koritschoner, N MEDLINE |1991 |138 |Am J Pathol 11257 Kuhn, C |HCAPLUS 12996 |Genes Dev |11||1997 Kuo, C HCAPLUS 11986 Science 1997 |277 Kuo, C

| HCAPLUS 1235 Gene 1195 11997 Lalazar, A HCAPLUS 1575 |Cell 11993 172 Leuther, K |Nucleic Acids Res | HCAPLUS 13796 1995 123 Maruyama, I

|Mol Cell Biol | HCAPLUS |13 12776 1993 Miller, I |EMBO J 11609 11995 | 4 Miller, J

| HCAPLUS 1371 **IScience** |1989 |245 Mitchell, P | HCAPLUS |1995 |375 |316 Nature Nuez, B | HCAPLUS |Genomics |1998 |48 1143 Onyango, P

| HCAPLUS Nature |318 |1995 1375 Perkins, A |Proc Natl Acad Sci U| 195 19500 11998 Ratzin, V

|Molecular Cloning:A 11989 Sambrook, J |1986 |47 |Cell 11025 Schur, R |1986 |271 J Biol Chem 120009 Shields, J

| J Biol Chem 1272 118504 11988 Shields, J | HCAPLUS |Nucleic Acids Res 1796 |1998 126 Shields, J

|Nucleic Acids Res 11527 11993 |21 Sogo, K |Mol Cell Biol | HCAPLUS 1442 118 11998 Watanabe, T **IHCAPLUS** [J Biol Chem |1998 |273 1026

COPYRIGHT 2002 ACS ANSWER 10 OF 37 HCAPLUS L57

**HCAPLUS** 1998:294709 AN

129:145538 DN

Yet, S

Berg, J Chen, X

Gill, G

Chomczynski, P

Garret-Sinha, L

Crossley, M El Rouby, S Friedman, S

Helios, a T cell-restricted Ikaros family member that quantitatively  ${ t TI}$ associates with Ikaros at centromeric heterochromatin. [Erratum to document cited in CA129:23889]

Hahm, Kyungmin; Cobb, Bradley S.; McCarty, Aaron S.; Brown, Karen E.; AU Klug, Christopher A.; Lee, Robert; Akashi, Koichi; Weissman, Irving L.; Fisher, Amanda G.; Smale, Stephen T.

Howard Hughes Medical Institute, Molecular Biology Institute, and CS Department of Microbiology and Immunology, UCLA School of Medicine, Los Angeles, CA, 90095-1662, USA

Genes Dev. (1998), 12(8), 1240 SO CODEN: GEDEEP; ISSN: 0890-9369

Cold Spring Harbor Laboratory Press PB

Journal DT

English LA

The name of Irving L. Weissman was spelled incorrectly in the Table of ABContents of this issue.

207870-86-8 ΙT

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence) (amino acid sequence of limiting regulatory subunit of Ikaros; cdna sequence of mouse helios T cell-restricted Ikaros family member that quant. assocs. with Ikaros at centromeric heterochromatin (Erratum))

L57 ANSWER 11 OF 37 HCAPLUS COPYRIGHT 2002 ACS 1998:289907 HCAPLUS AN





- DN 129:23889
- TI Helios, a T cell-restricted Ikaros family member that quantitatively associates with Ikaros at centromeric heterochromatin
- AU Hahm, Kyungmin; Cobb, Bradley S.; McCarty, Aaron S.; Brown, Karen E.; Klug, Christopher A.; Lee, Robert; Akashi, Koichi; Weissman, Irving L.; Fisher, Amanda G.; Smale, Stephen T.
- CS Howard Hughes Medical Institute, Molecular Biology Institute, and Department of Microbiology and Immunology, UCLA School of Medicine, Los Angeles, CA, 90095-1662, USA
- SO Genes Dev. (1998), 12(6), 782-796 CODEN: GEDEEP; ISSN: 0890-9369
- PB Cold Spring Harbor Laboratory Press
- DT Journal
- LA English
- The Ikaros gene encodes multiple protein isoforms that contribute crit. AB functions during the development of lymphocytes and other hematopoietic cell types. The intracellular functions of Ikaros are not known, although recent studies have shown that Ikaros proteins colocalize with inactive genes and centromeric heterochromatin. In this study, Ikaros proteins were found to be components of highly stable complexes. The complexes from an immature T cell line were purified, revealing assocd. proteins of 70 and 30 kD. The p70 gene, named Helios, encodes two protein isoforms with zinc finger domains exhibiting considerable homol. to those within Ikaros proteins. Helios and Ikaros recognize similar DNA sequences and, when overexpressed, Helios assocs. indiscriminately with the various Ikaros isoforms. Although Ikaros is present in most hematopoietic cells, Helios was found primarily in T cells. The relevance of the Ikaros-Helios interaction in T cells is supported by the quant. assocn. of Helios with a fraction of the Ikaros. Interestingly, the Ikaros-Helios complexes localize to the centromeric regions of T cell nuclei, similar to the Ikaros localization previously obsd. in B cells. Unlike the B cell results, however, only a fraction of the Ikaros, presumably the fraction assocd. with Helios, exhibited centromeric localization in T cells. These results establish immunoaffinity chromatog. as a useful method for identifying Ikaros partners and suggest that Helios is a limiting regulatory subunit for Ikaros within centromeric heterochromatin.
- IT 207870-86-8

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence) (DNA-binding, amino acid sequence of limiting regulatory subunit of Ikaros; cdna sequence of mouse helios T cell-restricted Ikaros family member that quant. assocs. with Ikaros at centromeric heterochromatin)

- L57 ANSWER 12 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1998:78103 HCAPLUS
- DN 128:227538
- TI End effects in DNA recognition by zinc finger arrays
- AU Choo, Yen
- CS Medical Research Council Laboratory of Molecular Biology, Cambridge, CB2 2QH, UK
- SO Nucleic Acids Res. (1998), 26(2), 554-557 CODEN: NARHAD; ISSN: 0305-1048
- PB Oxford University Press
- DT Journal
- LA English
- The paradigmatic DNA binding domain from the transcription factor Zif268 contains three zinc finger modules in tandem repeat. When bound to their cognate DNA site the fingers read out the sequence of one DNA strand by making a linear series of successive base contacts. It is shown that the base-specific protein-DNA contacts made from the ends of the Zif268 three-finger array contribute less to the stability of the intermol. complex than do structurally equiv. contacts from more central regions of the DNA binding domain.

The effect is akin to the end fraying obsd. in duplex nucleic acid mols.

204594-56-9 IT

RL: PEP (Physical, engineering or chemical process); PROC (Process) (amino acid sequence; end effects in DNA recognition by zinc finger arrays)

ANSWER 13 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57

1997:698566 HCAPLUS AN

128:30912 DN

Solution structure of the first three zinc fingers of TITFIIIA bound to the cognate DNA sequence: determinants of affinity and sequence specificity

Wuttke, Deborah S.; Foster, Mark P.; Case, David A.; Gottesfeld, Joel M.; ΑU Wright, Peter E.

Department of Molecular Biology and the Skaggs Institute for Chemical CS Biology, La Jolla, CA, 92037, USA

J. Mol. Biol. (1997), 273(1), 183-206 SO CODEN: JMOBAK; ISSN: 0022-2836

Academic PB

Journal DT

English LA

AB

The high resoln. soln. structure of a protein contg. the 3 N-terminal zinc fingers of Xenopus laevis transcription factor IIIA (TFIIIA) bound to its cognate DNA duplex was detd. by NMR spectroscopy. The protein, which is designated zf1-3, binds with all 3 fingers in the DNA major groove, with a no. of amino acids making base-specific contacts. The DNA structure is close to B-form. Although the mode of interaction of zf1-3 with DNA is similar to that of zif268 and other structurally characterized zinc finger complexes, the TFIIIA complex exhibits several novel features. zinc finger contacts 4-5 base-pairs and the repertoire of known base contact residues is extended to include a tryptophan at position +2 of the helix (finger 1) and arginine at position +10(finger 3). Sequence-specific base contacts are made over virtually the entire length of the finger 3 helix. Lysine and histidine side-chains involved in base recognition are dynamically disordered in the soln. structure; in the case of lysine, in particular, this could significantly decrease the entropic cost of DNA binding. The TGEKP(N) linker sequences, which are highly flexible in the unbound protein, adopt ordered conformations on DNA binding. The linkers appear to play an active structural role in stabilization of the protein-DNA complex. Substantial protein-protein contact surfaces are formed between adjacent fingers. As a consequence of these protein-protein interactions, the orientation of finger 1 in the major groove differs from that of the other fingers. Contributions to high affinity binding by zf1-3 come from both direct protein-DNA contacts and from indirect protein-protein interactions assocd. with structural organization of the linkers and formation of well-packed interfaces between adjacent zinc fingers in the DNA complex. The structures provide a mol. level explanation for the large body of footprinting and mutagenesis data available for the TFIIIA-DNA complex. 159575-54-9

ΙT

RL: BPR (Biological process); PRP (Properties); BIOL (Biological study); PROC (Process)

(soln. structure of the first three zinc fingers of TFIIIA bound to the cognate DNA sequence and determinants of affinity and sequence specificity)

L57 ANSWER 14 OF 37 HCAPLUS COPYRIGHT 2002 ACS

1997:643827 HCAPLUS AN

127:327914 DN





TI Synthesis and **DNA-binding** ability of Sp1 protein zinc finger domain and its peptidomimetics

Wang, Rui; Ni, Jingman; Hu, Xiaoyu; Ma, Yaping; Li, Xiangqun; Yang, Dingjian; Dong, Shouliang; Yang, Xiaowu; Pan, Xinfu

CS State Key Lab. Applied Organic Che,., Lanzhou Univ., Lanzhou, 730000, Peop. Rep. China

SO Sci. China, Ser. C: Life Sci. (1997), 40(5), 518-523 CODEN: SCCLFO; ISSN: 1006-9305

PB Science in China Press

DT Journal

ΑU

LA English

The second zinc finger fragment of Sp1 (Sp1-ZF2), its mutant (Sp1-ZF2/HT. E20.fwdarw.H, R23.fwdarw.T), and two mimic analogs (ZF20 and ZF15) were synthesized by stepwise solid phase technique. The CD spectra and UV-visible spectrum with CoCl2 indicated that the formation of zinc finger structure was affected not only by the hydrophobic amino acids but also by the change of the distance between Cys and His. Gel-retardation electrophoresis assays indicated that the Glu and Arg residues are very important for recognition. A single zinc finger like Sp1-ZF2 is able to bind DNA sequence specifically.

IT 197923-45-8P 197923-46-9P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (synthesis and DNA-binding ability of Spl protein zinc finger domain and peptidomimetics)

L57 ANSWER 15 OF 37 HCAPLUS COPYRIGHT 2002 ACS

AN 1997:245950 HCAPLUS

DN 126:302855

TI A novel human zinc finger protein that interacts with the core promoter element of a TATA box-less gene

AU Koritschoner, Nicolas P.; Bocco, Jose L.; Panzetta-Dutari, Graciela M.; Dumur, Catherine I.; Flury, Alfredo; Patrito, Luis C.

CS Departmento de Bioquimica Clinica, Facultad de Ciencias Quimicas, Universidad Nacional de Cordoba, Cordoba, Argent.

SO J. Biol. Chem. (1997), 272(14), 9573-9580 CODEN: JBCHA3; ISSN: 0021-9258

PB American Society for Biochemistry and Molecular Biology

DT Journal

English LAThe authors describe a novel human cDNA isolated by target site screening AΒ of a placental expression library, using as a probe, an essential element of a TATA box-less promoter corresponding to a pregnancy-specific glycoprotein gene. The cDNA encoded a predicted protein of 290 amino acids, designated core promoter-binding protein (CPBP), which has three zinc fingers (type Cys2-His2) at the end of its C-terminal domain, a serine/threonine-rich central region and an acidic domain lying within the N-terminal region. Addnl. sequence anal. and data base searches revealed that only the zinc finger domains are conserved (60-80% identity) in other transcription factors. In cotransfection assays, CPBP increased the transcription from a minimal promoter contg. its natural DNA-binding site. Moreover, a chimeric protein between CPBP and Gal4 DNA binding domain also increased the activity of an heterologous reporter gene contg. Gal4 DNA binding sites. The tissue distribution anal. of CPBP mRNA revealed that it is differentially expressed with an apparent enrichment in placental cells. The DNA binding and transcriptional activity of CPBP, in conjunction with its expression pattern, strongly suggests that this protein may participate in the regulation and/or maintenance of the basal expression of PSG and possibly other TATA box-less genes.

IT 189284-91-1

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(amino acid sequence; cDNA sequence of a novel human zinc finger protein that interacts with the core promoter element of a TATA box-less gene)

- L57 ANSWER 16 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1997:103680 HCAPLUS
- DN 126:182011
- TI Identification of amdX, a new Cys-2-His-2 (C2H2) zinc-finger gene involved in the regulation of the amdS gene of Aspergillus nidulans
- AU Murphy, Rachael L.; Andrianopoulos, Alex; Davis, Meryl A.; Hynes, Michael J.
- CS Department of Genetics, The University of Melbourne, Parkville, 3052, Australia
- SO Mol. Microbiol. (1997), 23(3), 591-602 CODEN: MOMIEE; ISSN: 0950-382X
- PB Blackwell
- DT Journal
- LA English
- The acetamidase-encoding amdS gene of Aspergillus nidulans has been shown to be controlled by multiple regulatory genes. A new gene, amdX, involved in amdS regulation was identified during the characterization of a translocation affecting amdS control. The amdX gene is predicted to encode a 1150-amino-acid poly-peptide which contains two Cys-2-His-2 (C2H2) zinc

finger DNA-binding motifs. Insertional

inactivation of amdX and the phenotypes of transformants contg. multiple copies of the amdX gene show that it is an activator of amdS expression. A fusion protein contg. the AmdX zinc fingers

was found to bind to sequences in the 5' region of amdS which overlap binding sites for the CreA and AmdA regulatory proteins. Evidence is presented for AmdX acting at these sites in vivo.

IT 187414-36-4

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(amino acid sequence; identification of amdX, a new Cys-2-His-2 (C2H2) zinc-

finger gene involved in regulation of amdS gene of Aspergillus nidulans)

- L57 ANSWER 17 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1996:424019 HCAPLUS
- DN 125:134177
- TI A zinc finger gene from Onchocerca volvulus encodes a protein with a functional signal peptide and an unusual Ser-His finger motif
- AU Holst, Corinna; Zipfel, Peter F.
- CS Dep. Molecular Biol., Bernhard Nocht Inst. for Tropical Medicine, Hamburg, 20359, Germany
- SO J. Biol. Chem. (1996), 271(28), 16725-16733 CODEN: JBCHA3; ISSN: 0021-9258
- DT Journal
- LA English
- The filarial parasite Onchocerca volvulus is the causative agent of river blindness. In order to identify genes potentially involved in parasite development we cloned a zinc finger-encoding gene from this species. The ovzf-1 gene represents one member of a family of related zinc finger genes. The predicted ovzf-1 translation product of 447 amino acids includes a hydrophobic signal peptide, which is followed by 13 contiguous finger motifs. The domains of fingers II-XIII display several conserved amino acids and a typical Krueppel-like Cys2-His2 motif. The first finger domain has the two conserved Cys residues replaced with Ser residues; however, it includes all addnl. amino acids typical of

zinc finger domains. The N-terminal domain functions as a signal peptide, as it directs secretion of a reporter protein and a truncated Ovzf protein. Expression of an Ovzf protein via the secretory pathway was also confirmed by demonstrating attachment of N-linked carbohydrates to the recombinant protein. Although the recombinant Ovzf protein also includes a signal peptide, immunofluorescence analyses localize it inside a specific compartment of the infected insect cell. Expression of ovzf mRNA is developmentally regulated; no specific transcript is detected in adult female worms but in the infective L3. Identification of a secreted protein that might function in modulating gene expression of host cells provides an interesting tool for the study of parasite-host interaction on a biochem. and mol. level.

IT180033-42-5

> RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)

(amino acid sequence; comparison of Onchocerca volvulus ovzf-1 and ovzf-2 gene products)

IT180033-43-6

> RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)

(amino acid sequence; isolation and characterization of zincfinger encoding gene ovzf-1 in Onchocerca volvulus)

- L57 ANSWER 18 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1996:419878 HCAPLUS
- DN125:106854
- A single amino acid determines the specificity for the target sequence of TItwo zinc-finger proteins in plants
- Takatsuji, Hiroshi AU
- CS National Inst. Agrobiol. Resources, Tsukuba, 305, Japan
- Biochem. Biophys. Res. Commun. (1996), 224(1), 219-223 SO CODEN: BBRCA9; ISSN: 0006-291X
- DT Journal
- English LA
- The EPF family is a group of DNA-binding AB proteins with two canonical Cys2/His2 zinc-finger motifs in Petunia. These proteins are unique in terms of structure in that (i) the two zinc fingers are sepd. by spacers of various lengths and (ii) the sequence QALGGH is strongly conserved in the zinc-finger motifs of members of the family. In this study, domain-swapping and site-directed mutagenesis expts. with two members of the protein family. EPF2-5 and EPF2-7, which have different target sequences, revealed that only a single amino acid in the second zinc finger is responsible for the difference in target specificity. The postion of this amino acid is different from those of determinants of target-sequence specificity in other zinc-finger proteins. Thus, the EPF family recognizes target sequences in a unique manner, together with the recognition of spacings in the target sequence that we demonstrated recently.

179339-47-0 IT

RL: BPR (Biological process); PRP (Properties); BIOL (Biological study); PROC (Process)

(second zinc finger of EPF2-5, DNA

binding specificity of; single amino acid dets. specificity for target sequence of two zinc-finger proteins in plants)

- L57 ANSWER 19 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- ΑN 1996:320586 HCAPLUS
- 125:27275 DN
- TIA new Cys2/His2 zinc finger gene,

rKr1, expressed in oligodendrocytes and neurons

- Pott, Uwe; Colello, Raymond J.; Schwab, Martin E. AU
- CS Brain Research Institute, University of Zurich, August Forel-Strasse 1, Zurich, CH-8029, Switz.

```
SO Mol. Brain Res. (1996), 38(1), 109-121
CODEN: MBREE4; ISSN: 0169-328X
```

DT Journal

LA English

The myelination of nerve fibers is essential for the function of the AB vertebrate nervous system as a prerequisite for fast saltatory conduction of action potentials. In the central nervous system (CNS), myelin is produced by oligodendrocytes. In order to identify gene regulatory proteins involved in the differentiation of this glial cell type or in the expression of myelin-specific genes, we have constructed a cDNA library from a highly enriched population of rat oligodendrocytes and screened this library for members of the Krueppel family of Cys2 /His2 zinc finger proteins. One of the identified clones, named rKr1, encodes a novel protein of 650 amino acids which contains 12 carboxy-terminal zinc finger domains and an amino-terminal acidic domain. On Northern blots, a single rKr1 mRNA of 4.3 kb is detected. This message is present in all adult rat tissues tested, with the highest levels found in the CNS and testis. In situ hybridization on the P15 brain revealed that the transcript is expressed in differentiated oligodendrocytes and in subtypes of neurons. Particularly high message levels are found in motor neurons of the brainstem and the spinal cord. The modular structure of the rKrl protein, in which a potential DNA binding region (the zinc fingers) is combined with a putative activation domain (the acidic region), suggests a function as sequence-specific transcriptional activator.

IT 177773-66-9

RL: PRP (Properties)

(amino acid sequence; Cys2/His2 zinc finger gene rKrl expressed in oligodendrocytes and neurons)

L57 ANSWER 20 OF 37 HCAPLUS COPYRIGHT 2002 ACS

AN 1996:182634 HCAPLUS

DN 124:252560

TI Purification and characterization of the DNA-binding domain of BTEB, a GC box-binding transcription factor, expressed in Escherichia coli

AU Kikuchi, Yasuo; Sogawa, Kazuhiro; Watanabe, Nobuaki; Kobayashi, Akira; Fujii-Kuriyama, Yoshiaki

CS Dep. Chem., Graduate Sch. Sci., Tohoku Univ., Sendai, 980, Japan

SO J. Biochem. (Tokyo) (1996), 119(2), 309-13 CODEN: JOBIAO; ISSN: 0021-924X

DT Journal

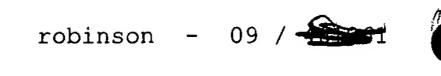
LA English

BTEB is a GC-binding protein that regulates the ABtranscription of genes with a single GC-box or tandemly repeated GC-boxes in the promoter. The DNA-binding domain of BTEB consists of 3 contiguous Cys2-His2 zinc finger motifs and short segments adjacent to their N- and C-terminal sides. The truncated BTEB (residues 120-244) contg. the DNA-binding domain was expressed in Escherichia coli and purified to homogeneity under denaturing conditions. DNAbinding activity of the BTEB was regenerated by refolding in the presence of Zn2+. The efficiency in regeneration was 70 .+-. 10%, and the dissocn. const. (Kd) of the DNA-complex was 4 .+-. 2 nM. Co2+ also regenerated the DNA-binding affinity of BTEB, albeit with less efficiency than Zn2+. Co-BTEB showed a slightly lower affinity to the specific DNA than Zn-BTEB. Refolding in the presence of Cd2+ resulted in an extremely low efficiency in regeneration of the DNA-binding activity. Zn -BTEB is in a monomer state at concns. < 0.5 .mu.M, and forms a dimer in the concn. range of about 10-100 .mu.M.

IT 174883-39-7P

RL: BAC (Biological activity or effector, except adverse); PRP (Properties); PUR (Purification or recovery); BIOL (Biological study); PREP (Preparation)

(amino acid sequence; purifn. and characterization of DNA-binding



domain of BTEB, a GC box-binding transcription factor, expressed in Escherichia coli)

- L57 ANSWER 21 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1995:895545 HCAPLUS
- DN 124:47002
- A new Cys2/His2 zinc finger gene, ΤI

rKr2, is expressed in differentiated rat oligodendrocytes and encodes a protein with a functional repressor domain

- Pott, Uwe; Thiesen, Hans-Juergen; Colello, Raymond J.; Schwab, Martin E. AU
- Brain Res. Inst., Univ. Zurich, Zurich, Switz. CS
- J. Neurochem. (1995), 65(5), 1955-66 SO

CODEN: JONRA9; ISSN: 0022-3042

- Journal DT
- English  $_{
  m LA}$
- The function of the vertebrate nervous system is dependent on the proper ABmyelination of its fiber tracts. Myelin of the CNS is produced by oligodendrocytes. To identify gene regulatory proteins expressed in this particular glial cell type, cDNAs coding for Cys2/His2 zinc finger

proteins were isolated from a rat oligodendrocyte cDNA library. One clone, named rKr2 (rKr for rat Krueppel-type protein), encodes a protein with 19 C-terminal zinc

finger domains and an N-terminal Krueppel-assocd. box domain. This N-terminal domain of the rKr2 protein behaved as a strong transcriptional repressor module when fused to the DNA-

binding domain of yeast GAL4 and tested on an appropriate reporter construct. High levels of rKr2 mRNA in adult rat tissues were found only in the CNS and testis; in the CNS, the message was predominantly expressed in differentiated oligodendrocytes. The modular structure of the rKr2 protein (C-terminal DNA-binding domain,

N-terminal repressor module) and its expression pattern suggest that it acts as a sequence-specific transcriptional repressor in the myelin-producing glial cells of the CNS.

172020-95-0 172020-96-1 IT

> RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)

(amino acid sequence; Cys2/His2 zinc

finger gene rKr2 is expressed in differentiated rat oligodendrocytes and encodes a protein with a functional repressor domain)

- ANSWER 22 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57
- 1995:856117 HCAPLUS AN
- 123:248582 DN
- Derivatives of zinc finger nucleic acid-binding ΤI domains of transcription factors and their use in the modulation of gene expression
- Barbas, Carlos F., III; Gottesfeld, Joel M.; Wright, Peter E. IN
- Scripps Research Institute, USA PA
- PCT Int. Appl., 135 pp. SO

CODEN: PIXXD2

- Patent DT
- English LA

FAN.	CNT 2			
•	PATENT NO.	KIND DATE	APPLICATION NO.	DATE
ΡI	WO 9519431	A1 19950720	WO 1995-US829	19950118 <
	W: AU, CA,	FI, JP, NO, US		
	RW: AT, BE,	CH, DE, DK, ES, FR,	GB, GR, IE, IT, LU,	MC, NL, PT, SE
	CA 2181548	AA 19950720	CA 1995-2181548	19950118 <
	AU 9516865	A1 19950801	AU 1995-16865	19950118 <
	AU 704601	B2 19990429		
	EP 770129	A1 19970502	EP 1995-908614	19950118 <
	R: AT, BE,	CH, DE, DK, ES, FR,	GB, GR, IE, IT, LI,	LU, MC, NL, PT, SE
	TP 09508019	T2 19970819	JP 1995-519231	

```
FI 1996-2879
    FI 9602879
                                                          19960717 <--
                           19960917
                          19960918
                                          NO 1996-2991
                                                          19960717 <--
    NO 9602991
                      A
                                         US 1996-676318
    US 6242568
                                                          19961230 <--
                      B1 20010605
                      A 19940118 <--
PRAI US 1994-183119
    US 1994-312604
                      A
                           19940928 <--
    WO 1995-US829
                           19950118 <--
                      W
```

AB An assay for identification of novel transcription-modulating zinc finger-nucleotide binding polypeptides is described. These proteins are useful for inhibiting, activating or enhancing gene expression from a zinc finger-nucleotide binding motif contg. promoter or other transcriptional control element, as well as a structural gene or RNA sequence and so may be of therapeutic use. Novel zinc finger-nucleotide binding polypeptides are described. The assay measures the binding of a protein to zinc finger-binding site and so does not require detailed knowledge of the structure of the protein. A panel of randomly mutagenized genes for zinc finger proteins in a phage display library are panned against an array of zinc finger recognition sequences.

IT 169108-70-7 169108-73-0 169108-74-1 169108-76-3

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological study); PROC (Process); USES (Uses).

(amino acid sequence; derivs. of zinc finger

nucleic acid-binding domains of transcription factors and their use in modulation of gene expression)

IT 168971-84-4D, analogs, derivs.

RL: MSC (Miscellaneous)

(zinc finger linker peptide; derivs. of

zinc finger nucleic acid-

binding domains of transcription factors and their use in modulation of gene expression)

L57 ANSWER 23 OF 37 HCAPLUS COPYRIGHT 2002 ACS

AN 1995:825211 HCAPLUS

DN 124:2339

TI Isolation and characterization of a novel zinc-finger protein with transcriptional repressor activity

AU Williams, Amy J.; Khachigian, Levon M.; Shows, Thomas; Collins, Tucker

CS Vascular Res. Div., Brigham and Women's Hosp. and Harvard Med. Sch., Boston, MA, 02115, USA

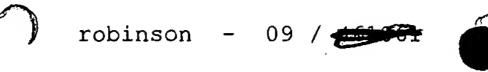
SO J. Biol. Chem. (1995), 270(38), 22143-52 CODEN: JBCHA3; ISSN: 0021-9258

DT Journal

LA English

To identify genes that can repress the expression of growth regulatory ABmols., a human fetal cDNA library was screened with a degenerate oligonucleotide that corresponds to the conserved stretch of 6 amino acids connecting successive zinc-finger regions in the Wilms' tumor suppressor/Egr-1 family of DNA-binding proteins. One clone, designated zinc-finger protein 174 (ZNF174), corresponds to a putative transcription factor with 3 zinc fingers and a novel finger -assocd. domain, designated the SCAN box. The three Cys2-His2-type zinc fingers are positioned at the C-terminus, whereas the 65-amino acid finger-assocd. SCAN box is located near the N-terminus. Chromosomal localization using somatic cell hybrid anal. and fluorescent in situ hybridization mapped the gene for ZNF174 to human chromosome 16p13.3. The 2.5-kb transcript from this gene is expressed in a variety of human organs, but most strongly in adult testis and ovary. Fusion of the upstream regulatory region of ZNF174 to the DNA-binding domain of GAL4 revealed that the gene could confer a repression function on the heterologous DNAbinding domain. ZNF174 selectively repressed reporter activity

driven by the platelet-derived growth factor-B chain and transforming



growth factor-.beta.l promoters and bound to **DNA** in a specific manner. This member of the C2H2-type **zinc-finger** family is a novel transcriptional repressor.

IT 171042-66-3

RL: BAC (Biological activity or effector, except adverse); PRP (Properties); BIOL (Biological study) (nucleotide sequence; isolation and characterization of zinc-finger protein ZNF174 with transcriptional repressor activity)

- L57 ANSWER 24 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1995:230394 HCAPLUS
- DN 122:49494
- TI Cooperative, non-specific binding of a zinc finger peptide to DNA
- AU Nedved, Michael L.; Moe, Gregory R.
- CS Department Chemistry Biochemistry, University Delaware, Newark, DE, 19716, USA
- SO Nucleic Acids Res. (1994), 22(22), 4705-11 CODEN: NARHAD; ISSN: 0305-1048
- DT Journal
- LA English

AB

- The DNA binding and structural properties of Xfin-31, a 25-amino acid zinc finger peptide, in the reduced, oxidized, and zinc complex forms, as well as the 14-residue helical segment of the zinc finger (residues 12-25) were compared using affinity coelectrophoresis (ACE) and CD spectroscopy. The zinc complex and oxidized peptides bind cooperatively to DNA, although the cooperativity factor, .omega., is >15-fold greater for the zinc complex. The reduced peptide in the absence of zinc and the helical segment do not bind cooperatively (.omega. = 1). Hence, the binding const. for singly contiguous sites (K.omega.) ranges over 100-fold for the various peptides even though the intrinsic binding consts. (K) are similar. An increase in binding order and affinity for the other forms of Xfin-31 is correlated with an increasing similarity of the CD spectrum to that of the Xfin-31 zinc complex. The surprising DNA binding activity of the oxidized peptide may result from hydrophobic interactions between the N-terminal loop formed by the Cys3-Cys6 disulfide bond and conserved hydrophobic residues in the C-terminal segment. Xfin-31 may be a particularly useful model for studying several poorly understood aspects of cooperative, nonspecific DNA binding since it is small, has a stable, well-defined structure, and structures of zinc fingers bound to DNA have been detd.
- IT 123658-20-8 123714-99-8 123714-99-8D,

zinc complex

RL: BPR (Biological process); BIOL (Biological study); PROC (Process) (cooperative, nonspecific binding of a zinc finger peptide to DNA)

- L57 ANSWER 25 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1995:195296 HCAPLUS
- DN 122:48105
- TI The zebrafish egrl gene encodes a highly conserved, zincfinger transcriptional regulator
- AU Drummond, Iain A.; Rohwer-Nutter, Patricia; Sukhatme, Vikas P.
- CS Department of Medicine, Harvard Medical School and Beth Israel Hospital, Boston, MA, 02215, USA
- SO DNA Cell Biol. (1994), 13(10), 1047-55 CODEN: DCEBE8; ISSN: 1044-5498
- DT Journal
- LA English
- AB The Egr family of transcriptional regulators comprise a group of genes which encode members of the Cys2-His2 glass of zinc-finger proteins. The authors have isolated a zebrafish egrl homolog by screening a zebrafish genomic library

with a mouse Egrl zinc finger probe. Southern blotting indicated the existence of a single zebrafish ergl gene and, as in higher vertebrates, the presence of related members of a larger gene family. Sequence anal. of the zebrafish ergl coding region revealed a high level of homol. to the mouse, rat, and human Egrl genes with the notable exception of a polymorphic, triplet nucleotide repeat sequence in the region coding for the amino terminus of the Egrl protein. The predicted DNA-binding, zinc-

finger domain protein sequence was strictly conserved. The 5' region of the zebrafish egrl gene contained a variety of transcription factor binding sites, also present in the mouse gene, for serum response factor, CREB and c-Ets. The zebrafish egrl transcript was approx. 3.4 kb in size and was expressed in adult zebrafish brain and muscle RNA, a pattern of expression similar to that obsd. in The potential for zebrafish egrl to function as a transcriptional regular was tested by constructing an expression vector contg. zebrafish egr1 coding sequences under the control of a cytomegalovirus promoter. This construct was found to activate transcription of a reporter plasmid bearing multiple Egrl binding sites when transiently cotransfected into mouse 3T3 cells. These results indicate that the structure, regulation, and function of the Egr1 gene have been highly conserved during vertebrate evolution and suggest an important role for this gene in growth and development.

159868-47-0 IT

RL: PRP (Properties)

(amino acid sequence; zebrafish egrl gene encodes highly conserved zinc-finger transcriptional regulator involved in development)

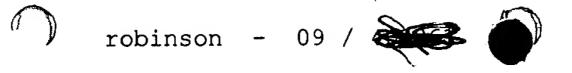
- ANSWER 26 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57
- 1995:127012 HCAPLUS AN
- 122:25024 DN
- The zebrafish egrl gene encodes a highly conserved, zinc-TIfinger transcriptional regulator
- Drummond, Iain A.; Rohwer-Nutter, Patricia; Sukhatme, Vikas P. ΑU
- Dep. Med., Harvard Med. Sch. and Beth Israel Hosp., Boston, MA, 02215, USA CS
- DNA Cell Biol. (1994), 13(9), 953-61 SO CODEN: DCEBE8; ISSN: 1044-5498
- Journal DT
- English LA
- The Egr family of transcriptional regulators comprises a group fo genes AB that encode members of the Cys2-His2 class of

zinc finger proteins. A zebrafish egrl

homolog was isolated by screening a zebrafish genomic library with a mouse egrl zinc finger probe. Southern blotting indicated the existence of single zebrafish egrl gene and, as in higher vertebrates, the presence of related members of a larger gene family. Sequence anal. of the zebrafish egrl coding region revealed a high level of homol. to the mouse, rat, and human egrl genes with the notable exception of a polymorphic, triplet and nucleotide repeat sequence in the region coding for the N-terminus of the egrl protein. The predicted

DNA-binding, zinc finger domain

protein sequence was strictly conserved. The 5' region of the zebrafish egrl gene contained a variety of transcription factor binding sites, also present in the mouse gene, for serum response factor, CREB, and c-ets. The zebrafish egrl transcript was .apprx.3.4 kb size and was expressed in adult zebrafish brain and muscle RNA, a pattern of expression similar to that obsd. in mice. The potential for zebrafish egrl to function as a transcriptional regulator was tested by constructing an expression vector contg. zebrafish egrl coding sequences under the control of a cytomegalovirus promoter. This construct activated transcription of a receptor plasmid bearing multiple egrl binding sites when transiently cotransfected into mouse 3T3 cells. The results indicate that the structure, regulation, and function of the egrl gene have been highly conserved during vertebrate evolution and suggest an important role for this gene in growth and development.



#### IT 159868-47-0

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(amino acid sequence; sequence and expression of the highly conserved zebrafish egr1 gene transcriptional regulator)

L57 ANSWER 27 OF 37 HCAPLUS COPYRIGHT 2002 ACS

AN 1994:673509 HCAPLUS

DN 121:273509

- TI Multiple products from the shavenbaby-ovo gene region of Drosophila melanogaster: relationship to genetic complexity
- AU Garfinkel, Mark D.; Wang, Jhy; Liang, Yuanping; Mahowald, Anthony P.
- CS Dep. Mol. Genet. Cell Biol., Univ. Chicago, Chicago, IL, 60637, USA
- SO Mol. Cell. Biol. (1994), 14(10), 6809-18 CODEN: MCEBD4; ISSN: 0270-7306
- DT Journal
- LA English
- The Drosophila melanogaster shavenbaby (svb)-ovo gene region is a complex ABlocus, contg. two distinct but comutable genetic functions. Ov is required for survival and differentiation of female germ line cells and plays a role in germ line sex detn. In contrast, svb is required in both male and female embryos for the prodn. of epidermal locomotor and sensory Sequences required for the two genetic functions are partially overlapping. Ovo corresponds to a previously described germ line-dependent 5.0-kb poly(A) + mRNA that first appears in the germarium and accumulates in nurse cells during oogenesis. The 5.0-kb mRNA is stored in the egg, but it is rapidly lost in the embryos except for its continued presence in the germ line precursor pole cells. The ovo mRNA predicts a 1028-amino-acid 110.6-kDa protein homologous with transcription factors. The authors have identified an embryonic mRNA, 7.1 kb in length, that contains exons partially overlapping those of the 5.0-kb poly(A) + mRNA. The spatial distribution of this newly discovered transcript during midembryogenesis suggests that it corresponds to the svb function. The arrangement of exons common to the 5.0- and 7.1-kb mRNAs suggests that the Ovo and Svb proteins share DNAbinding specificity conferred by four Cys2-His2 zinc finger motifs but differ functionally in their capacity to interact with other components of the transcription machinery.

IT 158857-29-5
RL: PRP (Properties)

(amino acid sequence; multiple products from shavenbaby-ovo gene region of Drosophila melanogaster and relationship to genetic complexity)

- L57 ANSWER 28 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1994:185581 HCAPLUS
- DN 120:185581
- TI Independence of metal binding between tandem Cys2His2 zinc finger domains
- AU Krizek, Beth Allyn; Zawadzke, Laura E.; Berg, Jeremy M.
- CS Dep. Chem., Johns Hopkins Univ., Baltimore, MD, 21218, USA
- SO Protein Sci. (1993), 2(8), 1313-9 CODEN: PRCIEI; ISSN: 0961-8368
- DT Journal
- LA English
- AB Most Cys2His2 zinc finger proteins contain tandem arrays of metal binding domains. The tandem nature of these arrays suggests that metal binding by these domains may not be independent but rather that metal binding may occur in a cooperative manner. This is esp. true in light of the crystal structure of a three zinc finger array bound to DNA that revealed several types of interactions between domains. To address this question, peptides contg. two tandem domains have been prepd. While metal binding studies do show that the two finger peptide has a metal ion affinity about threefold higher than that for a single domain peptide with the same sequence, addnl. studies reveal that this behavior is due to increased

single site affinities in the context of the two domain **peptide** rather than to cooperativity. These studies indicate that domains of this type are independent of one another with regard to metal **binding**, at least in the absence of **DNA**. This observation has implications with regard to the question of whether the activities of **proteins** of this class might be modulated by available **zinc** concns.

IT 133551-05-0

RL: BIOL (Biological study)
(cobalt binding to, in tandem zinc finger domain metal binding modeling)

L57 ANSWER 29 OF 37 HCAPLUS COPYRIGHT 2002 ACS

AN 1993:663879 HCAPLUS

DN 119:263879

TI The ht.beta. gene encodes a novel CACCC box-binding protein that regulates T-cell receptor gene expression

AU Wang, Yu Kang; Kobori, Joan A.; Hood, Leroy

CS Div. Biol., California Inst. Technol., Pasadena, CA, 91125, USA

SO Mol. Cell. Biol. (1993), 13(9), 5691-701 CODEN: MCEBD4; ISSN: 0270-7306

DT Journal

LA English

A gene encoding a novel CACCC box-binding protein that AB binds to the promoter region of the human T-cell receptor (TCR) V.beta.8.1 gene and the mouse TCR .alpha. gene silencer has been cloned. This gene, termed ht.beta., contains 4 zinc fingers of the class Cys2-X12-His2 that may be responsible for DNA binding and a highly neg. charged region that defines a putative transcriptional activation domain. Anal. of the expression of ht.beta. mRNA revealed similar expression levels and patterns in various cells The bacterially expressed ht.beta. protein can bind to the CACCC box in both the human TCR V.beta.8.1 gene promoter and the mouse TCR .alpha. gene silencer. The CACCC box is essential for efficient transcription of the V.beta.8.1 promoter. Cotransfection with an ht.beta. expression plasmid and a reporter vector indicated that ht.beta. can activate human transcription. Ht.beta. also is able to counteract the silencing effect of the mouse TCR .alpha. gene silencer. The CACCC box has been found in almost all V.beta.8.1 gene subfamily members and in both TCR .alpha. and .beta. gene enhancers in humans and mice. These results suggest that the CACCC box-binding protein may have an important regulatory function of TCR gene expression in .alpha..beta. T cells vs. .gamma..delta. T cells.

IT 151442-11-4

RL: PRP (Properties)
(amino acid sequence of and T-cell receptor gene expression regulation by)

L57 ANSWER 30 OF 37 HCAPLUS COPYRIGHT 2002 ACS

AN 1993:511528 HCAPLUS

DN 119:111528

TI Adjacent zinc-finger motifs in multiple zinc
-finger peptides from SWI5 form structurally independent,
flexibly linked domains

AU Nakaseko, Yukinobu; Neuhaus, David; Klug, Aaron; Rhodes, Daniela

CS Lab. Mol. Biol., MRC, Cambridge, CB2 2QH, UK,

SO J. Mol. Biol. (1992), 228(2), 619-36 CODEN: JMOBAK; ISSN: 0022-2836

DT Journal

LA English





Peptides contg. either one, two or three of the three AB zinc-finger motifs from the yeast transcription factor SWI5 have been prepd. by expression in Escherichia coli. The DNA binding characteristics of these peptides were investigated, and a two-dimensional NMR study undertaken to establish the three-dimensional structures of the two-finger peptide The peptide contq. fingers 1 and 2 binds sequence specifically to two thirds of the DNA binding site recognized either by intact SWI5 or by the isolated threefinger peptide, and hence has the correct tertiary fold for DNA recognition. These results also establish the polarity of DNA binding, since the N-terminal two fingers of SWI5 bind to the 5' end of the DNA binding site. Mild proteolysis of the three-finger peptide using trypsin results in a small no. of discrete products, which is consistent with the presence of three structured mini-domains. Nearly complete NMR signal assignments were obtained for two peptides contg. finger 2 alone or fingers 1 + 2. Comparison of two-dimensional spectra of these peptides and others clearly shows that the NOE enhancements and chem. neighboring fingers. This clearly indicates that adjacent zincfinger domains are structurally independent in these peptides from SWI5. However, there must be some steric limitations on the possible relative orientations of the fingers , and to establish limits for these a set of structures for the peptide contg. fingers 1 + 2 was calcd. using the YASAP simulated annealing protocol in conjunction with NMR-based constraints. A more detailed description of the three-dimensional structures of finger 1 and finger 2, and their relationship to other previously detd. structures of single zinc-fingers, is given in the accompanying paper. 128086-74-8 128087-09-2 146836-66-0 IT 146836-68-2 RL: BIOL (Biological study) (soln. structure and stability and DNA-binding properties of, as yeast SWI5 transcription factor zinc-finger motif model) 149348-55-0 149348-56-1 IT RL: BAC (Biological activity or effector, except adverse); PRP (Properties); BIOL (Biological study) (soln. structure and stability and DNA-binding properties of, as yeast SWI5 transcription factor zinc-finger motif model) ANSWER 31 OF 37 HCAPLUS COPYRIGHT 2002 ACS L571993:444590 HCAPLUS ANDN119:44590 Use of a zinc-finger consensus sequence ŢΙ framework and specificity rules to design specific DNA binding proteins Desjarlais, John R.; Berg, Jeremy M. AU Thomas C. Jenkins Dep. Biophys., Johns Hopkins Univ., Baltimore, MD, CS 21218, USA Proc. Natl. Acad. Sci. U. S. A. (1993), 90(6), 2256-60 SO CODEN: PNASA6; ISSN: 0027-8424

DT Journal LA English

DNA-binding specificities were designed. The design strategy combines a consensus zinc-finger framework sequence with previously characterized recognition regions such that the specificity of each protein is predictable. The 1st protein consists of 3 identical zinc fingers, each of which was expected to recognize the subsite GCG. This protein binds specifically to the sequence 5'-GCG-GCG-GCG-3' with a dissocn. const. of .apprxeq.11 .mu.M. The 2nd protein has 3 zinc fingers with different predicted preferred subsites. This protein binds to the predicted





recognition site 5'-GGG-GCG-GCT-3' with a dissocn. const. of 2 nM. Furthermore, selection expts. indicate that this is the optimal binding site. A permuted version of the 2nd protein was also constructed and shown to preferentially recognize the corresponding permuted site 5'-GGG-GCT-GCG-3' over the nonpermuted site. These results indicate that earlier observations on the specificity of zinc fingers can be extended to generalized zinc -finger structures and realize the use of zinc fingers for the design of site-specific DNAbinding proteins. This consensus-based design system provides a useful model system with which to study details of zinc -finger-DNA specificity. 148023-61-4, Protein QDR-RER-RHR (synthetic DNA -binding reduced) 148023-62-5, Protein RER-QDR-RHR (synthetic DNA-binding reduced) RL: BAC (Biological activity or effector, except adverse); PRP (Properties); BIOL (Biological study) (design and DNA binding by zinc-

finger motif in relation to)

148023-60-3, Protein RER-RER-RER (synthetic DNA

-binding reduced)

RL: BAC (Biological activity or effector, except adverse); PRP (Properties); BIOL (Biological study) (design and DNA binding by, zincfinger motif in relation to)

ANSWER 32 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57

1993:403327 HCAPLUS AN

119:3327 DN

IT

ΙT

- A novel human insulinoma-associated cDNA, IA-1, encodes a protein TIwith "zinc-finger" DNA-binding motifs
- Goto, Yasuhiro; De Silva, Mark G.; Toscani, Antonio; Prabhakar, Bellur S.; AU Notkins, Abner Louis; Lan, Michael S.
- Lab. Oral Med., Natl. Inst. Dent. Res., Bethesda, MD, 20892, USA CS
- J. Biol. Chem. (1992), 267(21), 15252-7 SO CODEN: JBCHA3; ISSN: 0021-9258

Journal DT

English LA

A subtraction library was constructed from human insulinoma (.beta. cell ABtumor) and glucagonoma (.alpha. cell tumor) cDNA phagemid libraries. Differential screening of 153 clones with end-labeled mRNAs from insulinoma, glucagonoma, and HeLa cells resulted in the isolation of a novel cDNA clone designated IA-1. This cDNA clone has a 2838-base pair sequence consisting of an open reading frame of 1530 nucleotides, which translated into a protein of 510 amino acids with a pI value of 9.1 and a mol. mass of 52,923 daltons. At the 3'-untranslated region there are seven ATTTA sequences between 2 polyadenylation signals (AATAAA). The IA-1 protein can be divided into 2 domains based upon the features of its amino acid sequence. The N-terminal domain of the deduced protein sequence (amino acids 1-250) has 4 classical pro-hormone dibasic conversion sites and an amidation signal sequence, Pro-Gly-Lys-Arg. The C-terminal domain (amino acids 251-510) contains five putative zinc-finger DNA-

binding motifs of the form X3-Cys-X2-4-Cys-X12-His-X3-4-His-X4 which has been described as a consensus sequence for members of the Cys2-His2 DNA-binding

protein class. Northern blot anal. revealed IA-1 mRNA in 5 of 5 human insulinoma and 3 of 3 murine insulinoma cell lines. Expression of this gene was undetectable in normal tissues. Addnl. tissue studies revealed that the message is expressed in several tumor cell lines of neuroendocrine origin including pheochromocytoma, medullary thyroid carcinoma, insulinoma, pituitary tumor, and small cell lung carcinoma. The restricted tissue distribution and unique sequence motifs suggest that this novel cDNA (clone may encode a protein assocd. with the transformation of neuroendocrine cells.





IT 147955-03-1, Protein (human clone IA-1 insulinoma-associated reduced)

RL: PRP (Properties)

(amino acid sequence of, complete)

- L57 ANSWER 33 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1993:185997 HCAPLUS
- DN 118:185997
- TI Characterization of a zinc finger DNAbinding protein expressed specifically in Petunia petals and seedlings
- AU Takatsuji, Hiroshi; Mori, Masaki; Benfey, Philip N.; Ren, Ling; Chua, Nam Hai
- CS Lab. Plant Mol. Biol., Rockefeller Univ., New York, NY, 10021, USA
- SO EMBO J. (1992), 11(1), 241-9 CODEN: EMJODG; ISSN: 0261-4189
- DT Journal
- LA English
- In Petunia, the expression of the 5-enolpyruvylshikimate-3-phosphate synthase gene (EPSPS) is tissue-specific and developmentally regulated. Nuclear exts. from Petunia petal contain a factor that interacts with the 5'-upstream region of EPSPS. DNase I footprinting expts. revealed 4 strong binding sites (EP1-EP4) and several weaker sites that appear to bind the same factor. A cDNA (EPF1) encoding a DNA-

binding protein that has similar binding

activity to that of the nuclear factor was isolated. The deduced amino acid sequences shows that the encoded protein, EPF1, contains 2 repeats of a Cys2/His2 zinc finger

motif. EPF1 and the factor detected in nuclear exts. differ in their mol. wt. and Zn2+ requirements. Nevertheless, Northern blot analyses showed that the expression pattern of EPF1 is remarkably similar to that of EPSPS. In addn., as detd. by translational fusion of the EPF1 upstream region to the .beta.-glucuronidase reporter gene, the cell- specific expression pattern of EPF1 in flower and seedling nearly identical to that of EPSPS. Taken together with the results of cis-element analyses, these observations suggest that EPF1 may be one of the factors involved in the activation of EPSPS.

IT **146989-67-5** 

RL: PRP (Properties)
(amino acid sequence of, complete)

- L57 ANSWER 34 OF 37 HCAPLUS COPYRIGHT 2002 ACS
- AN 1993:18158 HCAPLUS
- DN 118:18158
- TI Clone pAT 133 identifies a gene that encodes another human member of a class of growth factor-induced genes with almost identical zinc-finger domains
- AU Mueller, Hans Joachim; Skerka, Christine; Bialonski, Alexandra; Zipfel, Peter F.
- CS Bernhard Nocht Inst. Trop. Med., Hamburg, 2000/36, Germany
- SO Proc. Natl. Acad. Sci. U. S. A. (1991), 88(22), 10079-83 CODEN: PNASA6; ISSN: 0027-8424
- DT Journal
- LA English
- The structure and regulation of a gene represented by clone pAT 133, which is induced upon transition from a resting state (GO) through the early phase of the cell cycle (G1), is reported. The pAT 133 gene is immediately induced, with FOS-like kinetics, in human T cells and in fibroblasts. Primary structure anal. showed that the encoded protein contains 3 tandem Zn-finger sequences of the type

Cys2-Xaa12-His2. This Zn-finger

region, which is thought to bind DNA in a sequence-specific manner, was similar (>80% on the amino acid level) to 2 previously described transcription factors, pAT 225/EGR1 and pAT 591/EGR2. Except for the conserved Zn-finger domains, the amino acid sequences of the 3 proteins were distinct. This structural similarity suggested





that the pAT 133 gene encodes a transcription factor with a specific biol. function. Comparing the regulation of these related Znfinger-encoding genes showed coordinate induction upon mitogenic stimulation of resting T lymphocytes and of resting fibroblasts. However, upon transition from a proliferating (G1) to a resting state of the cell cycle the 3 genes were differently regulated. In human histiocytic U937 cells, mRNA of clone pAT 133 was constitutively expressed, whereas mRNA of pAT 225/EGR1 was induced upon induction of terminal differentiation. In contrast, mRNA representing pAT 591/EGR2 was not expressed in these cells. This difference in gene regulation suggests distinct biol. roles in the control of cell proliferation for the resp. proteins. 144997-43-3, Protein (human clone pAT133-17/pAT133-15 50.6-kilodalton reduced)

IT

RL: PRP (Properties)

(amino acid sequence of, complete)

ANSWER 35 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57

1992:145140 HCAPLUS AN

116:145140 DN

The early response gene NGFI-C encodes a zinc finger TItranscriptional activator and is a member of the GCGGGGGCG (GSG) element-binding protein family

Crosby, Seth D.; Puetz, John J.; Simburger, Kelli S.; Fahrner, Timothy J.; AU Milbrandt, Jeffrey

Sch. Med., Washington Univ., St. Louis, MO, 63110, USA CS

Mol. Cell. Biol. (1991), 11(8), 3835-41 SO CODEN: MCEBD4; ISSN: 0270-7306

Journal DT

English LA

A nerve growth factor-induced early-response gene encodes a Cys2 AB/His2 zinc finger protein NGFI-C. RNA blot anal. demonstrates that NGFI-C mRNA is induced within minutes of stimulation of PC12 cells by nerve growth factor and is similarly

activated in brain after a Metrazol-induced seizure. The cDNA sequence predicts a protein that contains three zinc

fingers which show striking homol. to the DNA-

binding regions of three previously reported zinc

finger proteins, NGFI-A, Krox-20, and the Wilms' tumor gene product. NGFI-C binds to the previously described DNA-

binding site of these three proteins, which is

GCGGGGGCG. Cotransfection expts. revealed that NGFI-C strongly activates transcription from this site in mammalian cells. The isolation of another early-response gene that encodes a member of the G(C/G)G or GSG elementbinding family should provide an opportunity to investigate the relative contributions of a family of transcription factors to the cell's

response to changes in its environment. 139874-91-2, Ribonucleic acid formation factor NGFI-C (rat clone ΙT pCMV-NGFI-C reduced)

RL: PRP (Properties)

(amino acid sequence of)

ANSWER 36 OF 37 HCAPLUS COPYRIGHT 2002 ACS L57

1990:492265 HCAPLUS ΑN

DN 113:92265

A DNA-binding protein containing two widely TI separated zinc finger motifs that recognize the same DNA sequence

Fan, Chen Ming; Maniatis, Tom ΑU

Dep. Biochem. Mol. Biol., Harvard Univ., Cambridge, MA, 02138, USA CS

Genes Dev. (1990), 4(1), 29-42 SO CODEN: GEDEEP; ISSN: 0890-9369

Journal DT

English LA

A full-length cDNA clone was isolated which encodes a protein AB (PRDII-BF1) that binds specifically to a pos. regulatory domain (PRDII) of the human IFN-.beta. gene promoter, and to a similar sequence present in a

no. of other promoters and enhancers. The sequence of this protein reveals two novel structural features. First, it is the largest sequence-specific DNA-binding protein reported to data (298 kD). Second, it contains two widely sepd. sets of C2-H2-type zinc fingers. Remarkably, each set of zinc fingers binds to the same DNA sequence motif with similar affinities and methylation interference patterns. Thus, this protein may act by binding simultaneously to reiterated copies of the same recognition sequence. Although the function of PRDII-BF1 is not known, the level of its mRNA is inducible by serum and virus, albeit with different kinetics. 128826-24-4 RL: PRP (Properties) (amino acid sequence of) ANSWER 37 OF 37 HCAPLUS COPYRIGHT 2002 ACS

L57

1989:226313 HCAPLUS AN

110:226313 DN

IT

- Molecular cloning, sequencing, and mapping of EGR2, a human early growth TIresponse gene encoding a protein with "zinc-binding finger" structure
- Joseph, Loren J.; Le Beau, Michelle M.; Jamieson, Gordon A., Jr.; Acharya, ΑU Sonia; Shows, Thomas B.; Rowley, Janet D.; Sukhatme, Vikas P.
- Howard Hughes Med. Inst., Univ. Chicago, Chicago, IL, 60637, USA CS
- Proc. Natl. Acad. Sci. U. S. A. (1988), 85(19), 7164-8 SO CODEN: PNASA6; ISSN: 0027-8424
- Journal DT
- English LA
- Early growth response gene-1 (Egr-1) is a mouse gene displaying fos-like AB induction kinetics in diverse cell types following mitogenic stimulation. Egr-1 encodes a protein with zinc-binding

finger structure. Zinc fingers are a

protein structural motif that serve as DNA-

binding domains in several transcriptional regulatory

proteins. Using low-stringency hybridization with an Egr-1 cDNA probe, a distinct human cDNA (designated EGR2 for early growth response gene-2) was identified, which is coregulated with EGR1 by fibroblast and lymphocyte mitogens; however, several stimuli that induce Egr-1 mRNA in PC12 (rat pheochromocytoma) cells do not induce Egr-2 mRNA. The cDNA sequence predicts a protein of 406 amino acids, including 3

tandem zinc fingers of the Cys2-His2

class. Strikingly, the deduced amino acid sequences of human EGR2 and mouse Egr-1 are 92% identical in the zinc finger region but show no similarity elsewhere. EGR2 Maps to human chromosome 10 at bands q21-22. Structure-function anal. of EGR2 and EGR1 proteins should provide insight into the mechanisms linking signal transduction and transcriptional regulation of gene expression.

120718-61-8, Protein (human clone ZAP2/ZAP8/ZAP32 gene EGR2 ITreduced) 120718-62-9, Protein (mouse clone Krox-20 gene EGR2 reduced)

RL: PRP (Properties)

(amino acid sequence of)

## => d all 38-49

- ANSWER 38 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L74
- AN1998:448099 BIOSIS
- PREV199800448099 DN
- Comprehensive DNA recognition through concerted interactions from adjacent TIzinc fingers.
- Isalan, Mark; Klug, Aaron; Choo, Yen (1) ΑU
- (1) MRC Lab. Molecular Biol., Hills Road, Cambridge CB2 2QH UK CS
- Biochemistry, (Sept. 1, 1998) Vol. 37, No. 35, pp. 12026-12033. SO ISSN: 0006-2960.
- Article DT

) robinson - 09 / ( )

English LA Zinc fingers are small DNA-binding AB modules noted for their occurrence in a large number of eukaryotic transcription factors, and their use in protein engineering. Although it was expected that zinc fingers can bind to a wide diversity of DNA sequences, previous studies using model zinc finger domains from Zif268 (and Sp1) have revealed a potential limitation to the DNA-binding specificity. For example, phage display selection of individual zinc fingers to recognize trinucleotide DNA subsites returned fingers that bound specifically only to triplets of the form GNN, i.e., triplets with guanine at the 5' end. Following our recently reported work (Isalan, M., Choo, Y., and Klug, A. (1997) Proc. Natl. Acad Sci. U.S.A. 94, 5617 -5621), we now show that this limitation can be overcome by the concerted randomization of certain amino acid positions in adjacent zinc fingers that specify overlapping DNA subsites. This illustrates an important mechanism underlying DNA recognition by arrays of zinc fingers, and points the way to improved strategies for the design of highly specific zinc finger proteins that bind any given nucleotide sequence. Genetics and Cytogenetics - General \*03502 CC Biochemical Studies - General \*10060 Major Concepts IT Molecular Genetics (Biochemistry and Molecular Biophysics) Chemicals & Biochemicals IT zinc finger: DNA-binding specificity; DNA: comprehensive recognition Miscellaneous Descriptors ΙT DNA-zinc finger interaction ANSWER 39 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L74 AN1997:295637 BIOSIS DN PREV199799594840 Synergy between adjacent zinc fingers in TIsequence-specific DNA recognition. Isalan, Mark; Choo, Yen (1); Klug, Aaron ΑU (1) Med. Res. Council Lab. Molecular Biol., Hills Rd., Cambridge CB2 2QH CS UK Proceedings of the National Academy of Sciences of the United States of SO America, (1997) Vol. 94, No. 11, pp. 5617-5621. ISSN: 0027-8424. Article DTEnglish LAZif268-like zinc fingers are generally regarded as ABindependent DNA-binding modules that each specify three base pairs in adjacent, but discrete, subsites. However, crystallographic evidence suggests that a contact also can occur from the second helical position of one finger to the subsite of the preceding finger. Here we show for the three-finger DNA-binding domain of the protein Zif268, and a panel of variants, that deleting the putative contact from finger 3 can affect the binding specificity for the 5' base in the adjoining triplet, which forms part of the binding site of finger 2. This finding demonstrates that Zif268-like zinc fingers can specify overlapping 4-bp subsites, and that sequence specificity at the boundary between subsites arises from synergy between adjacent fingers. This has important implications for the design and selection of zinc fingers with novel DNA binding specificities. Genetics and Cytogenetics - General \*03502 CC Comparative Biochemistry, General \*10010 Biochemical Methods - General \*10050

Biochemical Methods - General \*10050

Biochemical Studies - Nucleic Acids, Purines and Pyrimidines \*10062

Biochemical Studies - Proteins, Peptides and Amino Acids \*10064

Biochemical Studies - Minerals \*10069

Replication, Transcription, Translation \*10300

BC

IT

IT

AN

DN

TI

AU

CS

SO

DT

LA

AB

CC

BC

IT

IT

IT

Miscellaneous Descriptors

Biophysics - General Biophysical Techniques \*10504 Biophysics - Molecular Properties and Macromolecules \*10506 Metabolism - General Metabolism; Metabolic Pathways \*13002 Physiology and Biochemistry of Bacteria \*31000 Genetics of Bacteria and Viruses \*31500 \*06702 Enterobacteriaceae Major Concepts Biochemistry and Molecular Biophysics; Genetics; Metabolism; Methods and Techniques; Molecular Genetics (Biochemistry and Molecular Biophysics); Physiology Miscellaneous Descriptors ANALYTICAL METHOD; CRYSTALLOGRAPHY; DNA; DNA-BINDING MODULES; DNA-BINDING SPECIFICITIES; MOLECULAR GENETICS; SEQUENCE-SPECIFIC DNA RECOGNITION; STRAIN-TG1; TRANSCRIPTION FACTORS; ZINC FINGERS ORGN Super Taxa Enterobacteriaceae: Eubacteria, Bacteria ORGN Organism Name Escherichia coli (Enterobacteriaceae) ORGN Organism Superterms bacteria; eubacteria; microorganisms ANSWER 40 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L74 1998:32703 BIOSIS PREV199800032703 Promoter-specific activation of gene expression directed by bacteriophage-selected zinc fingers. Choo, Y. (1); Castellanos, A.; Garcia-Hernandez, B.; Sanchez-Garcia, I.; Klug, A. (1) Med. Res. Counc. Lab. Mol. Biol., Hills Road, Cambridge CB2 2QH UK Journal of Molecular Biology, (Oct. 31, 1997) Vol. 273, No. 3, pp. 525-532. ISSN: 0022-2836. Article English It has been shown that sequence-specific DNA-binding domains containing zinc fingers can be selected from libraries displayed on filamentous bacteriophage. The affinity and specificity of these peptides are well characterised in vitro, but few data are available to demonstrate specific DNA binding and discrimination between closely related DNA sequences in vivo. Transient transactivation assays were performed in mammalian cells, using expression plasmids which produce different amounts of a model transcription factor containing a phage-selected zinc finger DNA-binding domain, and reporter plasmids which carry systematic variations of the promoter sequence. When the intracellular concentration of the transcription factor was appropriate, activation of gene expression was absolutely dependent on a promoter having the same DNA sequence as that originally used to select the zinc finger domain by phage display. However, excessive intracellular concentrations of the transcription factor resulted in some less-specific DNA binding, leading to gene activation from similar promoters containing a maximum of two base changes. Thus, provided delivery is carefully controlled, highly specific control of gene expression in vivo can be achieved using artificial transcription factors containing phage-selected zinc finger DNA-binding domains. Genetics and Cytogenetics - General \*03502 Biochemical Studies - General \*10060 Genetics of Bacteria and Viruses \*31500 Bacterial Viruses - General 02700 Major Concepts Molecular Genetics (Biochemistry and Molecular Biophysics) Chemicals & Biochemicals artificial transcription factors; DNA binding domain

```
) robinson - 09 / 1 )
```

bacteriophage-selected zinc fingers; gene expression: promoter-specific activation; DNA binding ORGN Super Taxa Bacterial Viruses: Viruses, Microorganisms ORGN Organism Name filamentous bacteriophage (Bacterial Viruses) ORGN Organism Superterms Bacterial Viruses; Microorganisms; Viruses ANSWER 41 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L74 1995:519061 BIOSIS AN PREV199598533361  $\mathsf{DN}$ Gene regulatory proteins and their interaction with DNA. TIKlug, Aaron AU MRC Lab. Molecular Biology, Cambridge CB2 2QH UK CS Chambers, D. A. [Editor]. Annals of the New York Academy of Sciences, SO (1995) Vol. 758, pp. 143-160. Annals of the New York Academy of Sciences; DNA: The double helix: Perspective and prospective at forty years. Publisher: New York Academy of Sciences 2 East 63rd Street, New York, New York 10021, USA. Meeting Info.: Conference Chicago, Illinois, USA October 13-16, 1993 ISSN: 0077-8923. ISBN: 0-89766-906-1 (paper), 0-89766-905-3 (cloth). Book; Conference DTEnglish LAGeneral Biology - Symposia, Transactions and Proceedings of Conferences, CC Congresses, Review Annuals 00520 Genetics and Cytogenetics - General \*03502 Biochemical Studies - Nucleic Acids, Purines and Pyrimidines \*10062 Biochemical Studies - Proteins, Peptides and Amino Acids \*10064 Replication, Transcription, Translation \*10300 Biophysics - Molecular Properties and Macromolecules \*10506 Biophysics - Membrane Phenomena \*10508 Endocrine System - General \*17002 Major Concepts IT Biochemistry and Molecular Biophysics; Endocrine System (Chemical Coordination and Homeostasis); Genetics; Membranes (Cell Biology); Molecular Genetics (Biochemistry and Molecular Biophysics) Chemicals & Biochemicals IT LEUCINE Miscellaneous Descriptors IT BASIC LEUCINE ZIPPER; BETA-RIBBON MOTIF; BOOK CHAPTER; GENE EXPRESSION CONTROL; HELIX-TURN-HELIX MOTIF; HOMEODOMAIN PROTEIN; HORMONE RECEPTOR DNA-BINDING DOMAIN; MEETING PAPER; MOLECULAR MODEL; MOLECULAR RECOGNITION; TATA-BOX BINDING PROTEIN; THREE-DIMENSIONAL STRUCTURE; TRANSCRIPTION FACTOR; ZINC FINGER PROTEIN 61-90-5 (LEUCINE) RNANSWER 42 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. L74 1995:340395 BIOSIS ANPREV199598354695 DN Zinc fingers. TIKlug, Aaron (1); Schwabe, John W. R. ΑU (1) MRC Lab. Molecular Biology, Hills Road, Cambridge CB2 2QH UK CS FASEB Journal, (1995) Vol. 9, No. 8, pp. 597-604. SO ISSN: 0892-6638. General Review DTEnglish LAThe term zinc finger was first used to describe a AΒ 30-residue, repeated sequence motif found in an unusually abundant Xenopus transcription factor. It was proposed that each motif is folded around a central zinc ion to form an independent minidomain and that

30-residue, repeated sequence motif found in an unusually abundant kenopus transcription factor. It was proposed that each motif is folded around a central zinc ion to form an independent minidomain and that adjacent zinc fingers are combined as modules to make up a DNA-binding domain with the modules "gripping" the DNA (hence the term finger). We now know that these proposals were correct and that these DNA-binding

```
robinson - 09 /
```

motifs are found in many eukaryotic DNA-binding proteins. More recently, crystal structures of three different complexes between zinc finger domains and their target DNA binding sites have revealed a remarkably simple mode of interaction with DNA. The simplicity of the zinc finger structure, and of its interaction with DNA, is a very striking feature of this protein domain. After the discovery of the zinc finger motif, patterns of potential zinc ligands have been found in several other proteins, some of which also bind to DNA. Structural studies of these domains have revealed how zinc can stabilize quite diverse protein architectures. In total, 10 such small zinc-binding domains have been studied structurally. These form a diverse collection, but each in turn has been termed a zinc finger motif -although clearly what they have in common is only their zinc-binding property, which stabilizes an apparently autonomously folded unit. Genetics and Cytogenetics - General \*03502 Biochemical Studies - Nucleic Acids, Purines and Pyrimidines \*10062 Biochemical Studies - Proteins, Peptides and Amino Acids \*10064 Biochemical Studies - Minerals \*10069 Replication, Transcription, Translation \*10300 Biophysics - Molecular Properties and Macromolecules \*10506 Biophysics - Membrane Phenomena \*10508 Endocrine System - General \*17002 Virology - Animal Host Viruses \*33506 Retroviridae \*02623 Major Concepts Microbiology; Molecular Genetics (Biochemistry and Molecular Biophysics) Miscellaneous Descriptors DNA-BINDING MOTIF; LIM DOMAIN; NUCLEAR HORMONE

BC

ΙT

CC

Biochemistry and Molecular Biophysics; Endocrine System (Chemical Coordination and Homeostasis); Genetics; Membranes (Cell Biology);

IT

RECEPTOR; NUCLEOCAPSID PROTEIN; PROTEIN FOLDING; SECONDARY STRUCTURE; TRANSCRIPTION FACTOR; ZINC BINDING DOMAIN

ORGN Super Taxa

Retroviridae: Viruses

ORGN Organism Name

human immunodeficiency virus (Retroviridae)

ORGN Organism Superterms

microorganisms; viruses

- L74 ANSWER 43 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
- AN1995:419418 BIOSIS
- DNPREV199598433718
- Designing DNA-binding proteins in TIthe surface of filamentous phage.
- Choo, Yen; Klug, Aaron ΑU
- Med. Res. Counc., Lab. Mol. Biol., Hills Road, Cambridge CB2 2QH UK CS
- Current Opinion in Biotechnology, (1995) Vol. 6, No. 4, pp. 431-436. SO ISSN: 0958-1669.
- Article  $\mathsf{DT}$
- English LA
- Comparative Biochemistry, General \*10010 CC

Biochemical Methods - General \*10050

Biochemical Methods - Nucleic Acids, Purines and Pyrimidines \*10052

Biochemical Methods - Proteins, Peptides and Amino Acids \*10054

Biochemical Studies - General \*10060

Biochemical Studies - Nucleic Acids, Purines and Pyrimidines \*10062

Biochemical Studies - Proteins, Peptides and Amino Acids \*10064

Replication, Transcription, Translation \*10300

Biophysics - Molecular Properties and Macromolecules \*10506

Metabolism - General Metabolism; Metabolic Pathways \*13002

Metabolism - Proteins, Peptides and Amino Acids \*13012

```
robinson - 09 /
     Metabolism - Nucleic Acids, Purines and Pyrimidines *13014
     Genetics of Bacteria and Viruses *31500
     Microbiological Apparatus, Methods and Media *32000
     Virology - Bacteriophage *33504
     Bacterial Viruses - General
                                   *02700
     Major Concepts
        Biochemistry and Molecular Biophysics; Genetics; Metabolism; Methods
        and Techniques; Microbiology; Molecular Genetics (Biochemistry and
        Molecular Biophysics)
    Miscellaneous Descriptors
        BIOTECHNOLOGY; DNA PROTEIN INTERACTIONS; MOLECULAR EVOLUTION; MOLECULAR
        STRUCTURE; PHAGE DISPLAY; PROTEIN ENGINEERING
ORGN Super Taxa
        Bacterial Viruses - General: Viruses
ORGN Organism Name
       bacterial viruses (Bacterial Viruses - General)
ORGN Organism Superterms
       microorganisms; viruses
L74 ANSWER 44 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
    1995:34477 BIOSIS
    PREV199598048777
    Toward a code for the interactions of zinc fingers
    with DNA: Selection of randomized fingers displayed on phage.
    Choo, Yen; Klug, Aaron
    Med. Res. Council, Lab. Mol. Biol., Hills Rd., Cambridge CB2 2QH UK
    Proceedings of the National Academy of Sciences of the United States of
    America, (1994) Vol. 91, No. 23, pp. 11163-11167.
    ISSN: 0027-8424.
    Article
    English
    We have used two selection techniques to study sequence-specific
    DNA recognition by the zinc finger, a small,
    modular DNA-binding minidomain. We have chosen
    zinc fingers because they bind as independent modules
    and so can be linked together in a peptide designed to bind a
    predetermined DNA site. In this paper, we describe how a library
    of zinc ringers displayed on the surface of bacteriophage
    enables selection of fingers capable of binding to
    given DNA triplets. The amino acid sequences of selected ringers
    which bind the same triplet are compared to examine how sequence-specific
    DNA recognition occurs. Our results can be rationalized in terms
    of coded interactions between zinc ringers and DNA,
    involving base contacts from a few a-helical positions. In the paper
    following this one, we describe a complementary technique which confirms
    the identity of amino acids capable of DNA sequence
    discrimination from these positions.
    Genetics and Cytogenetics - General *03502
    Biochemical Methods - Proteins, Peptides and Amino Acids *10054
    Biochemical Studies - Nucleic Acids, Purines and Pyrimidines *10062
    Biochemical Studies - Proteins, Peptides and Amino Acids *10064
    Biophysics - Molecular Properties and Macromolecules *10506
    Genetics of Bacteria and Viruses *31500
    Virology - Bacteriophage *33504
    Bacterial Viruses - General *02700
    Major Concepts
       Biochemistry and Molecular Biophysics; Genetics; Methods and Techniques
    Miscellaneous Descriptors
```

BC

IT

GENETIC ENGINEERING; METHOD; SEQUENCE SPECIFIC RECOGNITION

ORGN Super Taxa

BC

IT

IT

AN DN

TI

ΑU

CS

SO

 $\mathsf{DT}$ 

LA

AΒ

CC

IT

Bacterial Viruses - General: Viruses

ORGN Organism Name

bacterial viruses (Bacterial Viruses - General)

ORGN Organism Superterms

microorganisms; viruses





- L74 ANSWER 45 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
- AN 1995:59043 BIOSIS
- DN PREV199598073343
- TI In vivo repression by a site-specific DNA-binding protein designed against an oncogenic sequence.
- AU Choo, Yen; Sanchez-Garcia, Isidro; Klug, Aaron
- CS Med. Res. Council, Lab. Mol. Biol., Hills Rd., Cambridge CB2 2QH UK
- SO Nature (London), (1994) Vol. 372, No. 6507, pp. 642-645. ISSN: 0028-0836.
- DT Article
- LA English
- AB A DNA-binding peptide comprising three zinc-fingers has been engineered to bind specifically to a unique nine-base-pair region of a BCR-ABL fusion oncogene In preference to the parent genomic sequences. Binding to the target oncogene in chromosomal DNA is possible In transformed cells in culture, and results in blockage of transcription. Consequently, murine cells rendered independent of growth factors by the action of the oncogene revert to factor dependence upon transient transfection with a vector expressing the peptide.
- CC Cytology and Cytochemistry Animal 02506
  Genetics and Cytogenetics Animal \*03506
  Biochemical Methods Proteins, Peptides and Amino Acids \*10054
  Biochemical Studies Nucleic Acids, Purines and Pyrimidines 10062
  Biochemical Studies Proteins, Peptides and Amino Acids \*10064
  Biophysics Molecular Properties and Macromolecules \*10506
  Neoplasms and Neoplastic Agents Biochemistry \*24006
- BC Muridae \*86375
- IT Major Concepts

Biochemistry and Molecular Biophysics; Genetics; Methods and Techniques; Tumor Biology

IT Miscellaneous Descriptors

BCR-ABL FUSION ONCOGENE; CANCER RESEARCH IMPLICATIONS; CHROMOSOMAL DNA; IN-VIVO BINDING; PROTEIN ENGINEERING

ORGN Super Taxa

Muridae: Rodentia, Mammalia, Vertebrata, Chordata, Animalia

ORGN Organism Name

mouse (Muridae)

ORGN Organism Superterms

animals; chordates; mammals; nonhuman mammals; nonhuman vertebrates; rodents; vertebrates

- L74 ANSWER 46 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
- AN 1993:432622 BIOSIS
- DN PREV199396087247
- TI A role in DNA binding for the linker sequences of the first three zinc fingers of TFIIIA.
- AU Choo, Yen; Klug, Aaron
- CS Med. Res. Council, Lab. Mol. Biol., Cambridge CB2 2QH UK
- SO Nucleic Acids Research, (1993) Vol. 21, No. 15, pp. 3341-3346. ISSN: 0305-1048.
- DT Article
- LA English
- AB Zinc fingers of the TFIIIA type are connected by short linker sequences between the structural units. Structural investigations by 2D NMR in solution and by X-ray crystallographic analyses of complexes with DNA point to a passive role for the linkers. We have therefore investigated the influence of the linker sequence on DNA binding using as a model the first three fingers of the protein TFIIIA. Insertion of certain heterologous linkers abolishes binding, and replacement of individual amino acids can reduce binding by factors of up to twenty-four.
- CC Genetics and Cytogenetics General \*03502
  Biochemical Studies Nucleic Acids, Purines and Pyrimidines \*10062
  Biochemical Studies Proteins, Peptides and Amino Acids \*10064
  Biochemical Studies Minerals 10069





Biophysics - Molecular Properties and Macromolecules \*10506

IT Major Concepts

Biochemistry and Molecular Biophysics; Genetics

IT Miscellaneous Descriptors

CONDENSED STRUCTURE FORMATION; DNA COLLAPSE; DNA TRANSFER METHOD; GENE TRANSFER; MOLECULE DELIVERY; POSITIVELY CHARGED LIPID BILAYER

- L74 ANSWER 47 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
- AN 1994:126855 BIOSIS
- DN PREV199497139855
- TI Co-chairman's remarks: Protein designs for the specific recognition of DNA.
- AU Klug, Aaron
- CS MRC Lab. Molecular Biol., Hills Road, Cambridge CB2 2QH UK
- SO Gene (Amsterdam), (1993) Vol. 135, No. 1-2, pp. 83-92. ISSN: 0378-1119.
- DT Article
- LA English
- The selective expression of a gene is achieved through the interaction of protein transcription factors with characteristic nucleotide sequences located in the regulatory region of the gene, which is usually distinct from the coding region. These proteins contain domains which bind specifically to the DNA sites (or response elements). Some general principles in the design of these DNA-binding domains are described, followed by examples of the different structural classes discovered so far and how they recognise their binding sites.
- CC Genetics and Cytogenetics General \*03502
  Biochemical Studies Nucleic Acids, Purines and Pyrimidines \*10062
  Biophysics Molecular Properties and Macromolecules \*10506
  Biophysics Membrane Phenomena \*10508
- BC \*00500
- IT Major Concepts

Biochemistry and Molecular Biophysics; Genetics; Membranes (Cell Biology)

IT Chemicals & Biochemicals

LEUCINE

IT Miscellaneous Descriptors

HELIX-TURN-HELIX; HOMEODOMAIN; HORMONE RECEPTORS; LEUCINE ZIPPER; MODULAR DESIGN; ZINC FINGERS

ORGN Organism Name

organisms - Unspecified)

- RN 61-90-5 (LEUCINE)
- L74 ANSWER 48 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
- AN 1993:95056 BIOSIS
- DN PREV199395050252
- Solution structures of two zinc-finger domains from SWI5 obtained using two-dimensional proton nuclear magnetic resonance spectroscopy: A zinc-finger structure with a third strand of beta-sheet.
- AU Neuhaus, David (1); Nakaseko, Yukinobu; Schwabe, John W. R. (1); Klug, Aaron (1)
- CS (1) MRC Lab. Molecular Biol., Hills Rd., Cambrdige CB2 2QH England SO Journal of Molecular Biology, (1992) Vol. 228, No. 2, pp. 637-651. ISSN: 0022-2836.
- DT Article
- LA English
- This paper describes the detailed three-dimensional structures of two zinc-finger domains from the yeast transcription factor SWI5, calculated using the results of the n.m.r. experiments described in the accompanying paper. The structure of finger 2 is essentially similar to those previously obtained by others for isolated, synthetic single zinc-finger domains in solution, and for the three zinc-finger peptide Zif268 in its crystalline complex with DNA. The N-terminal half of the





sequence forms a two-stranded, irregular beta-sheet containing both the metal-binding cysteine residues, while the remainder of the structure forms a helix. Approximately the first half of this helix is alpha-helical, whereas the C-terminal portion, including the two metal binding histidine residues, is 3-10 helical. Four invariant hydrophobic residues form a core to the structure. In contrast to all previously described structures of zinc-finger domains, finger 1 has an additional strand in the beta-sheet, formed by residues N-terminal to the formal start of the finger motif. This additional strands plays a role in stabilising the folded form of finger 1, since a two-finger peptide lacking the N-terminal residues showed folded structure in finger 2 but not in finger 1. Genetics and Cytogenetics - Plant \*03504 Biochemical Studies - Nucleic Acids, Purines and Pyrimidines 10062 Biochemical Studies - Proteins, Peptides and Amino Acids \*10064 Biophysics - Molecular Properties and Macromolecules \*10506 Fungi - Unspecified \*15000 Major Concepts Biochemistry and Molecular Biophysics; Genetics Miscellaneous Descriptors DNA BINDING PROTEIN; NMR; THREE DIMENSIONAL STRUCTURE; TRANSCRIPTION ACTIVATOR PROTEIN ORGN Super Taxa Fungi - Unspecified: Fungi, Plantae ORGN Organism Name fungi (Fungi - Unspecified); yeast (Fungi - Unspecified) ORGN Organism Superterms fungi; microorganisms; nonvascular plants; plants ANSWER 49 OF 49 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC. 1990:414201 BIOSIS MODE OF INTERACTION OF THE ZINC FINGER PROTEIN TFILLA WITH A 5S RNA GENE OF XENOPUS. CHURCHILL M E A; TULLIUS T D; KLUG A MEDICAL RES. COUNCIL LAB. MOL., HILLS RD., CAMBRIDGE CB2 2QH, ENGLAND. PROC NATL ACAD SCI U S A, (1990) 87 (14), 5528-5532. CODEN: PNASA6. ISSN: 0027-8424. BA; OLD English The zinc finger protein TFIIIA, a positive transcription factor of the 5S RNA gene, binds to an internal control region of 50 nucleotides. Two modes of binding have been considered for the TFIIIA-DNA complex, one of which has beef proposed on the basis of nuclease and chemical protection experiments and the other on model building. Since then, evidence has accumulated on the structures of individual components of the complex-for example, zinc finger polypeptides studied by NMR and a segment of the binding site analyzed by x-ray crystallography, but no high-resolution structural data on the TFIIIA-DNA complex itself are available. Probes used previously to study the TFIIIA-DNA complex do not react with every nucleotide of DNA, unlike hydroxyl radical, which cleaves DNA at every backbone position. We describe here the quantitative analysis of high-resolution hydroxyl radical footprints and suggest how the array of zinc fingers might interact with the double helix. Genetics and Cytogenetics - Animal \*03506 Biochemical Methods - Nucleic Acids, Purines and Pyrimidines 10052 Biochemical Methods - Proteins, Peptides and Amino Acids 10054 Biochemical Methods - Minerals 10059 Biochemical Studies - Nucleic Acids, Purines and Pyrimidines \*10062 Biochemical Studies - Proteins, Peptides and Amino Acids \*10064 Biochemical Studies - Minerals \*10069 Biophysics - Molecular Properties and Macromolecules \*10506

BCSalientia 85306

CC

BC

IT

IT

L74

AN

ΤI

ΑU

CS

SO

FS

LA

AΒ

CC

Miscellaneous Descriptors IT